

Biological Modeling of Neural Networks

Wulfram Gerstner

EPFL, Lausanne, Switzerland

TAs in 2020:

Martin Barry (head)

Valentin Schmutz

Georgios Iatropoulos

Christos Sourmpis

COURSE WEBPAGE:

Moodle

**Week 1: A first simple neuron model/
neurons and mathematics**

**Week 2: Hodgkin-Huxley models and
biophysical modeling**

**Week 3: Two-dimensional models and
phase plane analysis**

**Week 4: Two-dimensional models,
type I and type II models**

**Week 5,6: Associative Memory,
Hebb rule, Hopfield**

Week 7-10: Networks, cognition, learning

**Week 11,12: Noise models, noisy neurons
and coding**

**Week 13: Estimating neuron models for
coding and decoding: GLM**

Week x: Online video: Dendrites/Biophysics

Week xx: Density equations

LEARNING OUTCOMES

- Solve linear one-dimensional differential equations
- Analyze two-dimensional models in the phase plane
- Develop a simplified model by separation of time scales
- Analyze connected networks in the mean-field limit
- Formulate stochastic models of biological phenomena
- Formalize biological facts into mathematical models
- Prove stability and convergence
- Apply model concepts in simulations
- Predict outcome of dynamics
- Describe neuronal phenomena

Transversal skills

- Plan and carry out activities in a way which makes optimal use of available time and other resources.
- Collect data.
- Write a scientific or technical report.

Look at samples of past exams

Use a textbook,
(Use video lectures)
don't use slides (only)

miniproject

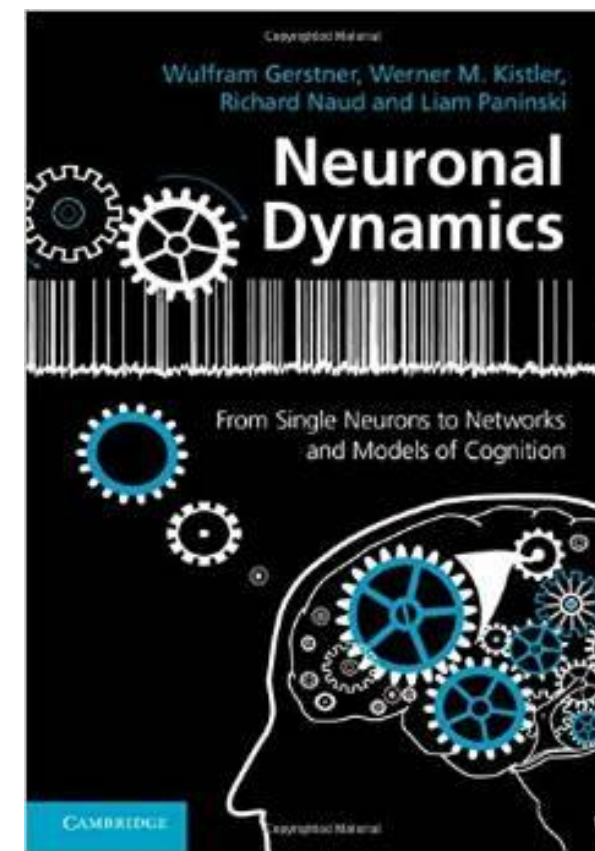
Biological Modeling of Neural Networks

Written Exam (70%)
+ miniproject (30%)

Miniproject consists of 3 extended computer exercises, of which you have to hand in 2 (first one is easier, recommended)

Textbook:

<http://neurondynamics.epfl.ch/>



Video:

<https://lcnwww.epfl.ch/gerstner/NeuronalDynamics-MOOC1.html>

<https://lcnwww.epfl.ch/gerstner/NeuronalDynamics-MOOC2.html>

Welcome back to EPFL!!

Today: Course in BS 170

BS 170 for the first week, but
INM 200 for the rest of the semester

Biological Modeling of Neural Networks



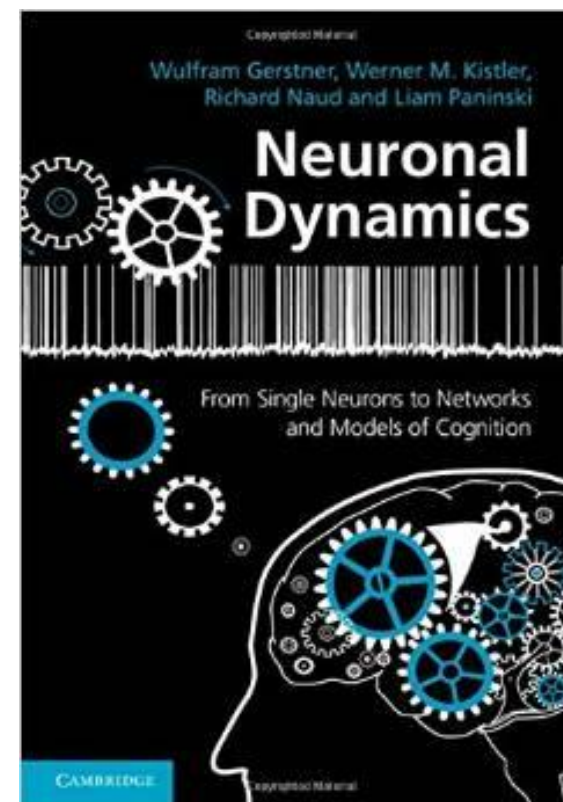
Week 1 – neurons and mathematics: a first simple neuron model

Wulfram Gerstner

EPFL, Lausanne, Switzerland

Reading for week 1:
NEURONAL DYNAMICS
- Ch. 1 (without 1.3.6 and 1.4)
- Ch. 5 (without 5.3.1)

Cambridge Univ. Press



1.1 Neurons and Synapses:

Overview

1.2 The Passive Membrane

- Linear circuit
- Dirac delta-function

1.3 Leaky Integrate-and-Fire Model

1.4 Generalized Integrate-and-Fire Model

1.5. Quality of Integrate-and-Fire Models

Biological Modeling of Neural Networks



1.1 Neurons and Synapses:

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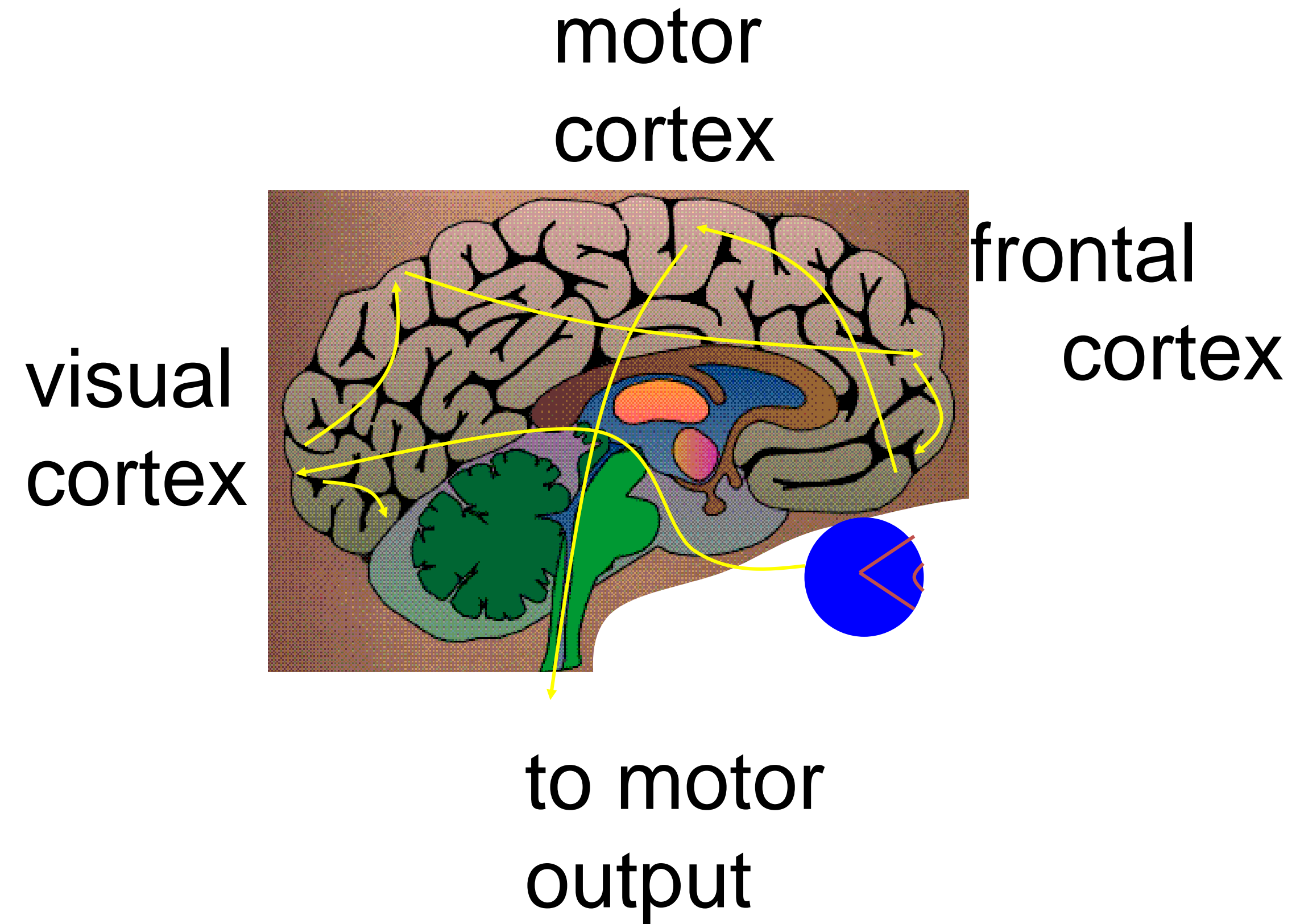
1.3 Leaky Integrate-and-Fire Model

1.4 Generalized Integrate-and-Fire Model

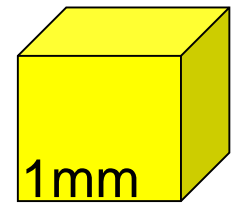
1.5. Quality of Integrate-and-Fire Models

Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

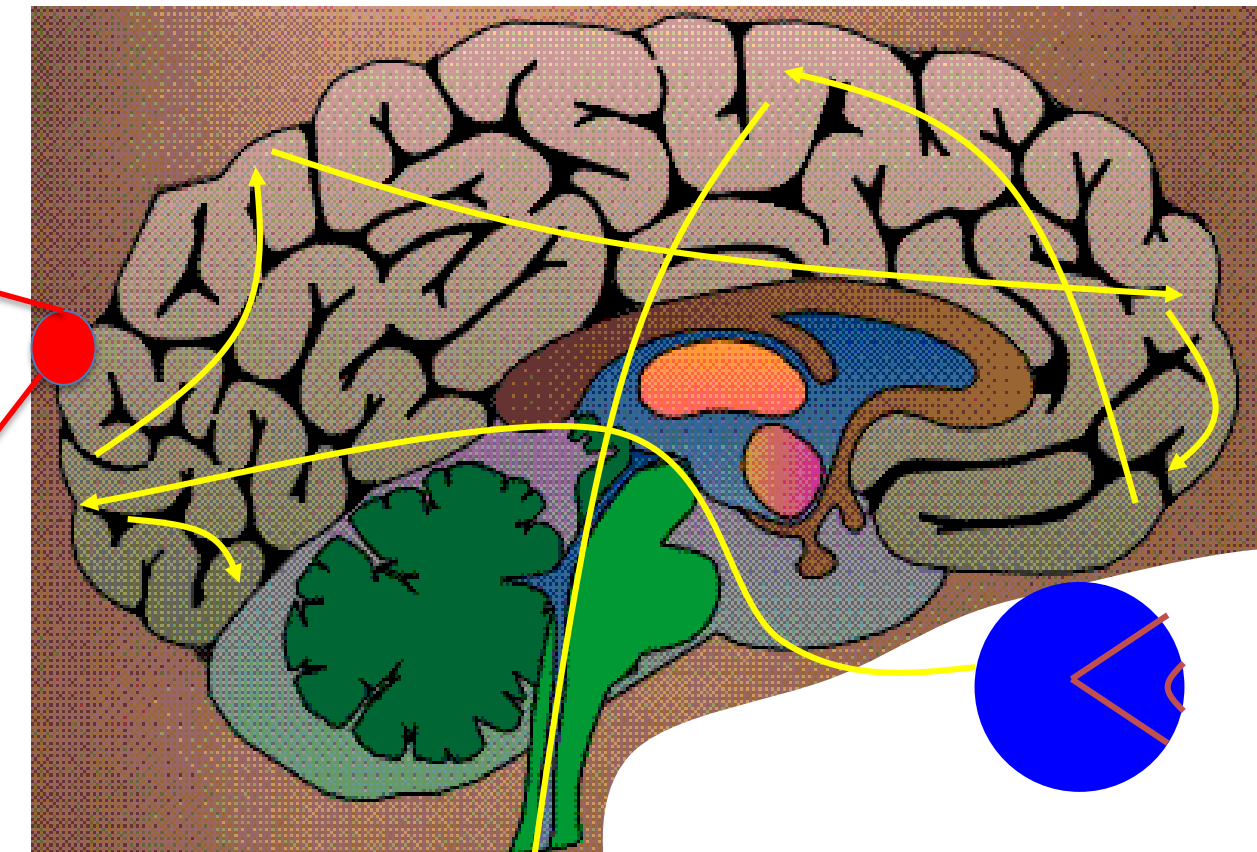
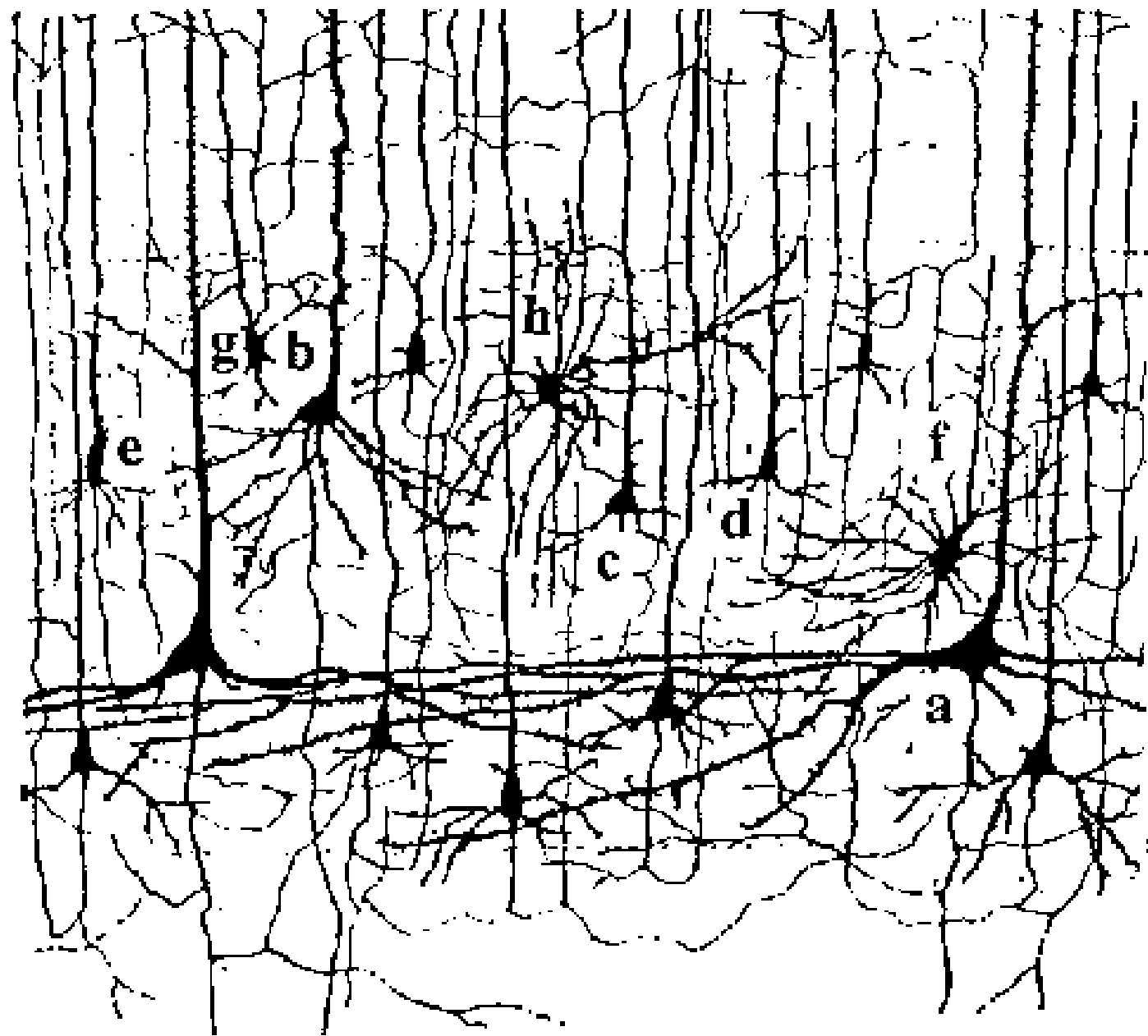
How do we recognize things?
Models of cognition
Weeks 5-10



Neuronal Dynamics – 1.1. Neurons and Synapses/Overview



10 000 neurons
3 km of wire

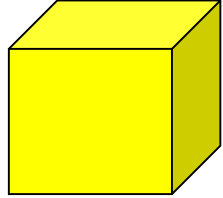


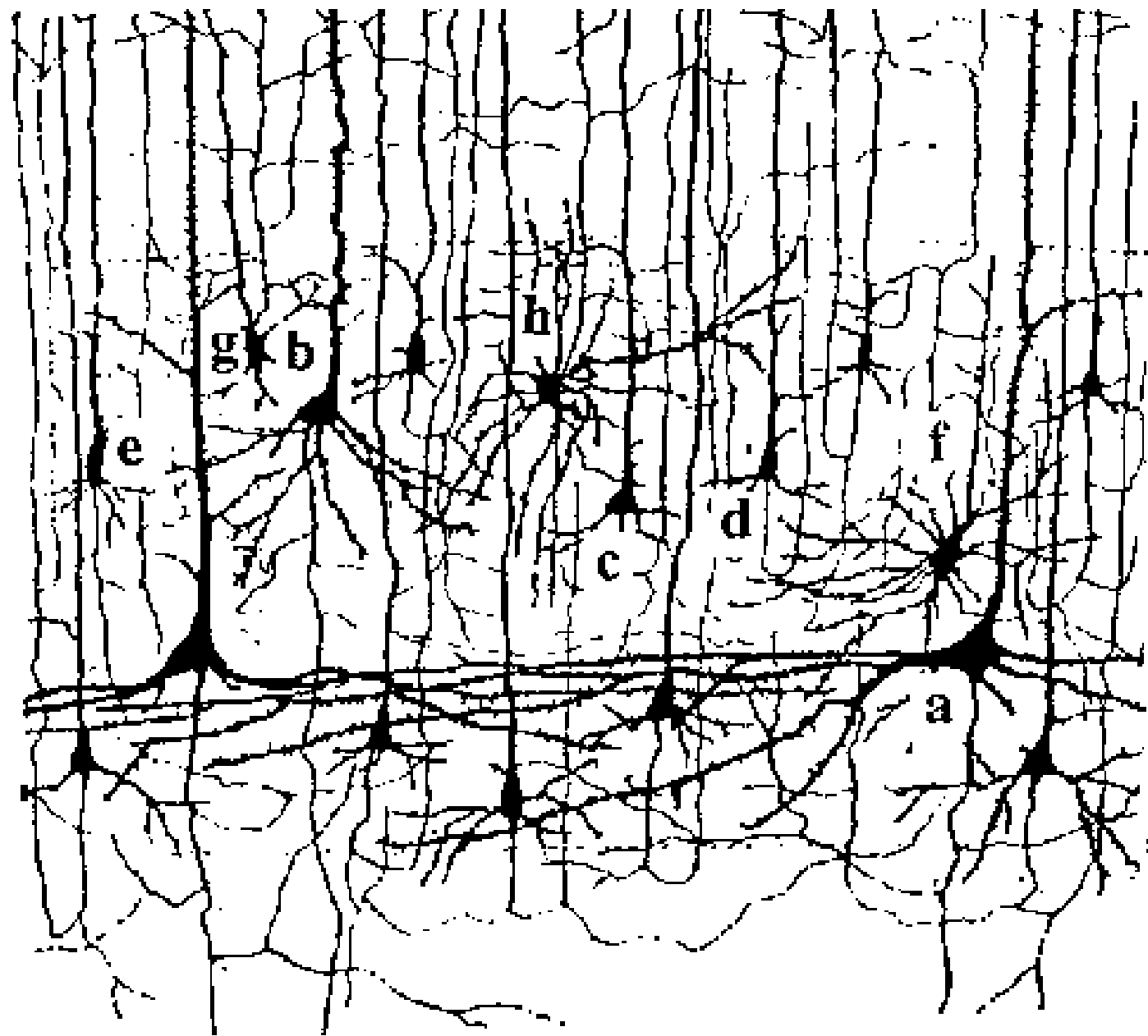
motor
cortex

frontal
cortex

to motor
output

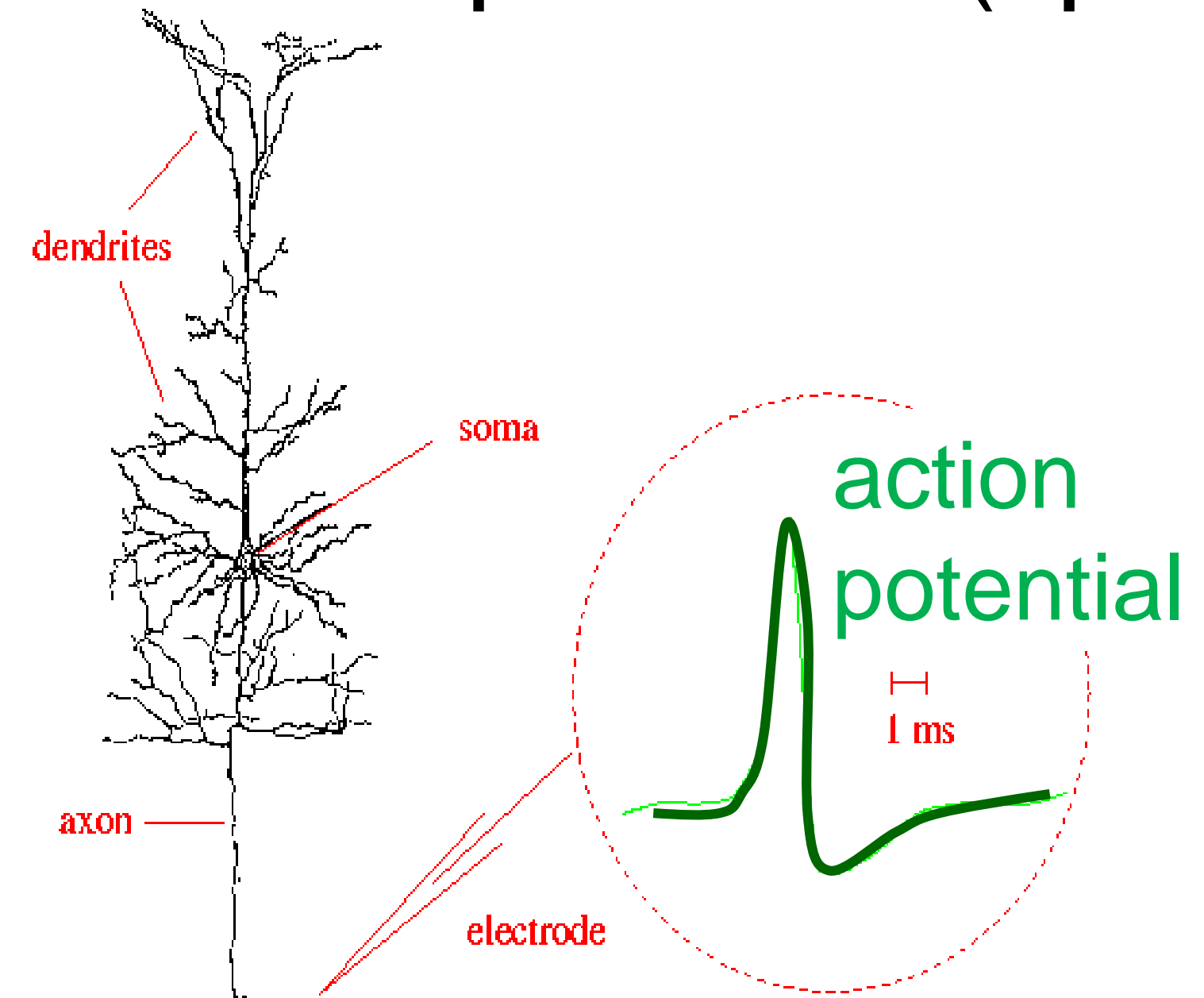
Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

 10 000 neurons
1mm 3 km of wire



Ramon y Cajal

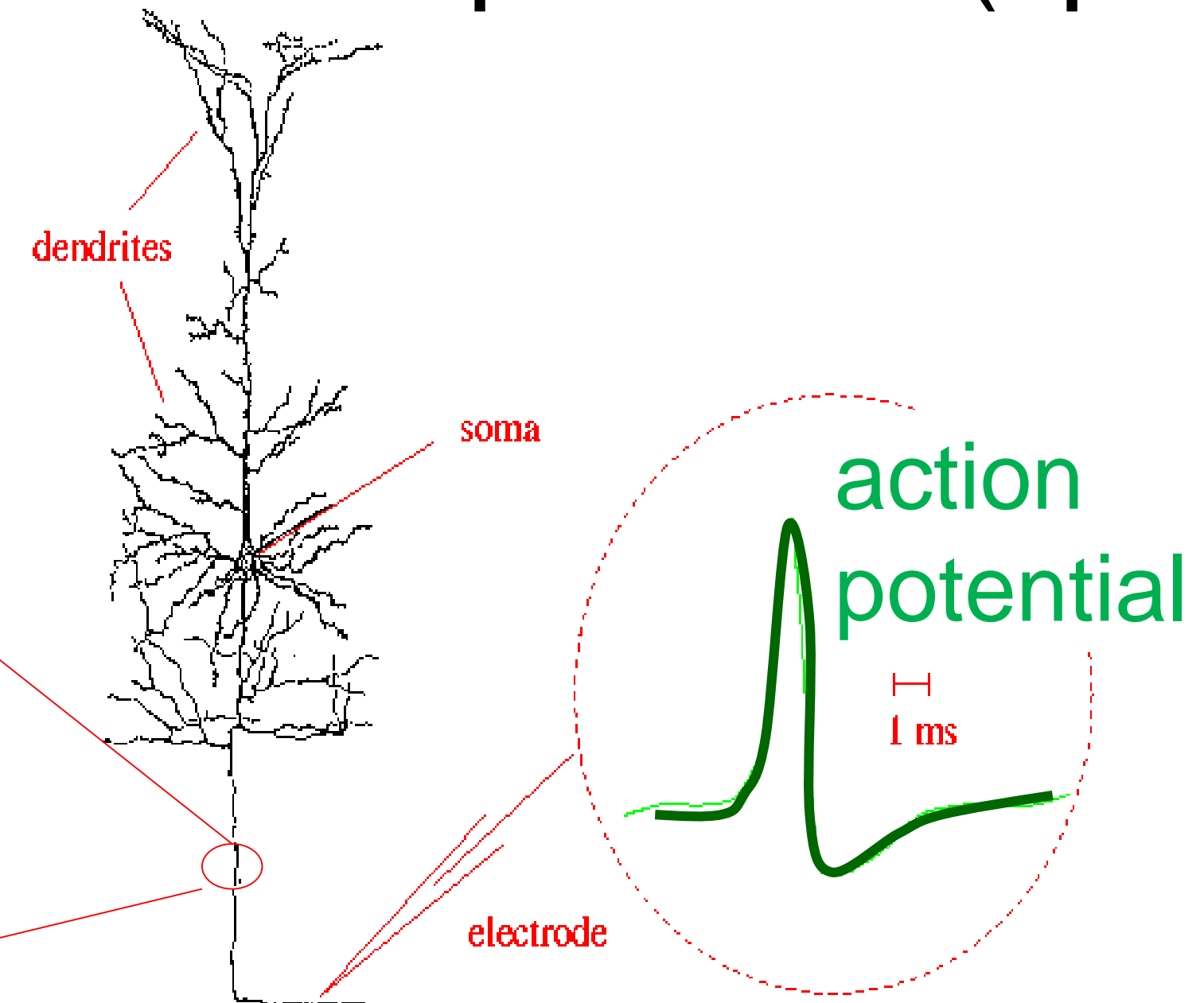
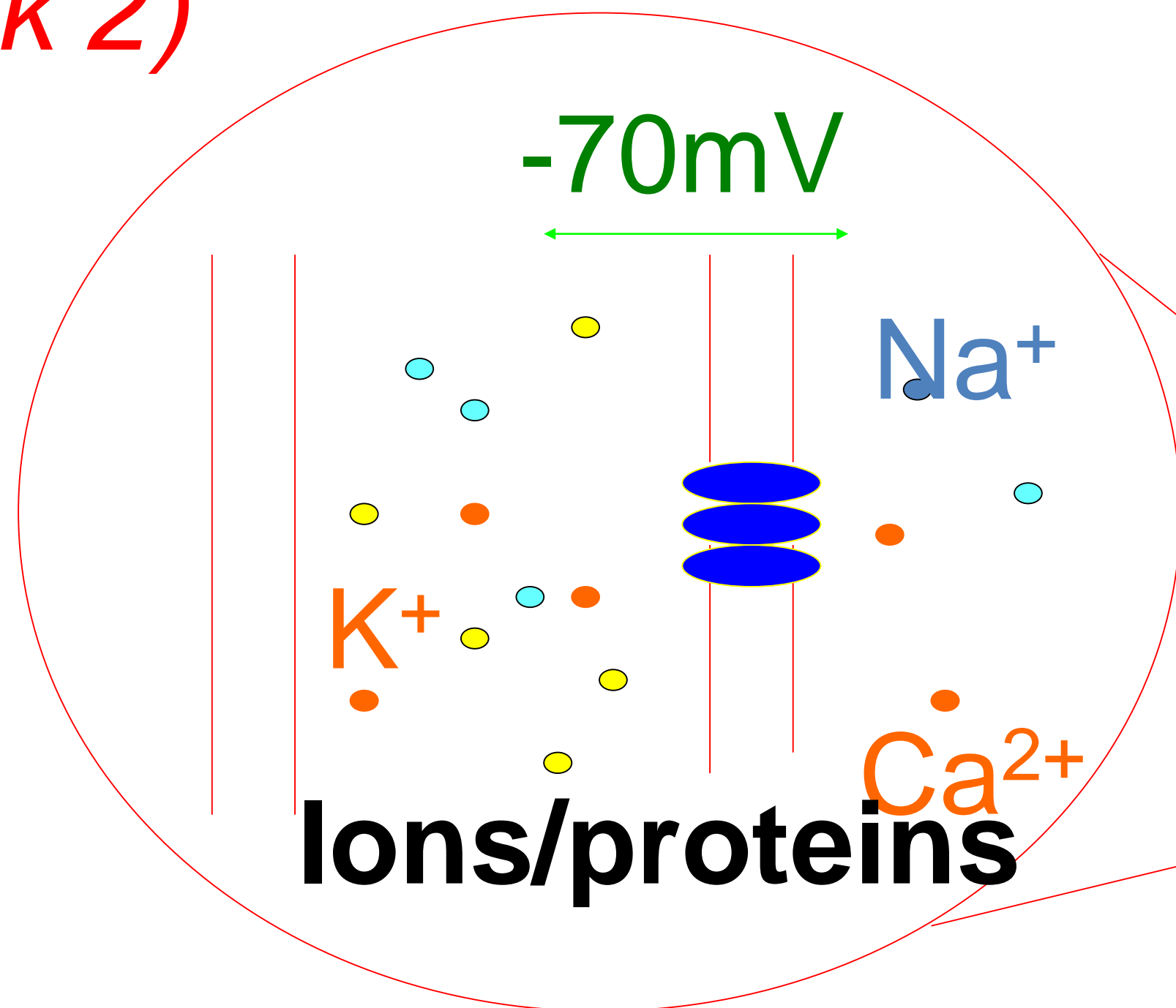
Signal:
action potential (spike)



Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

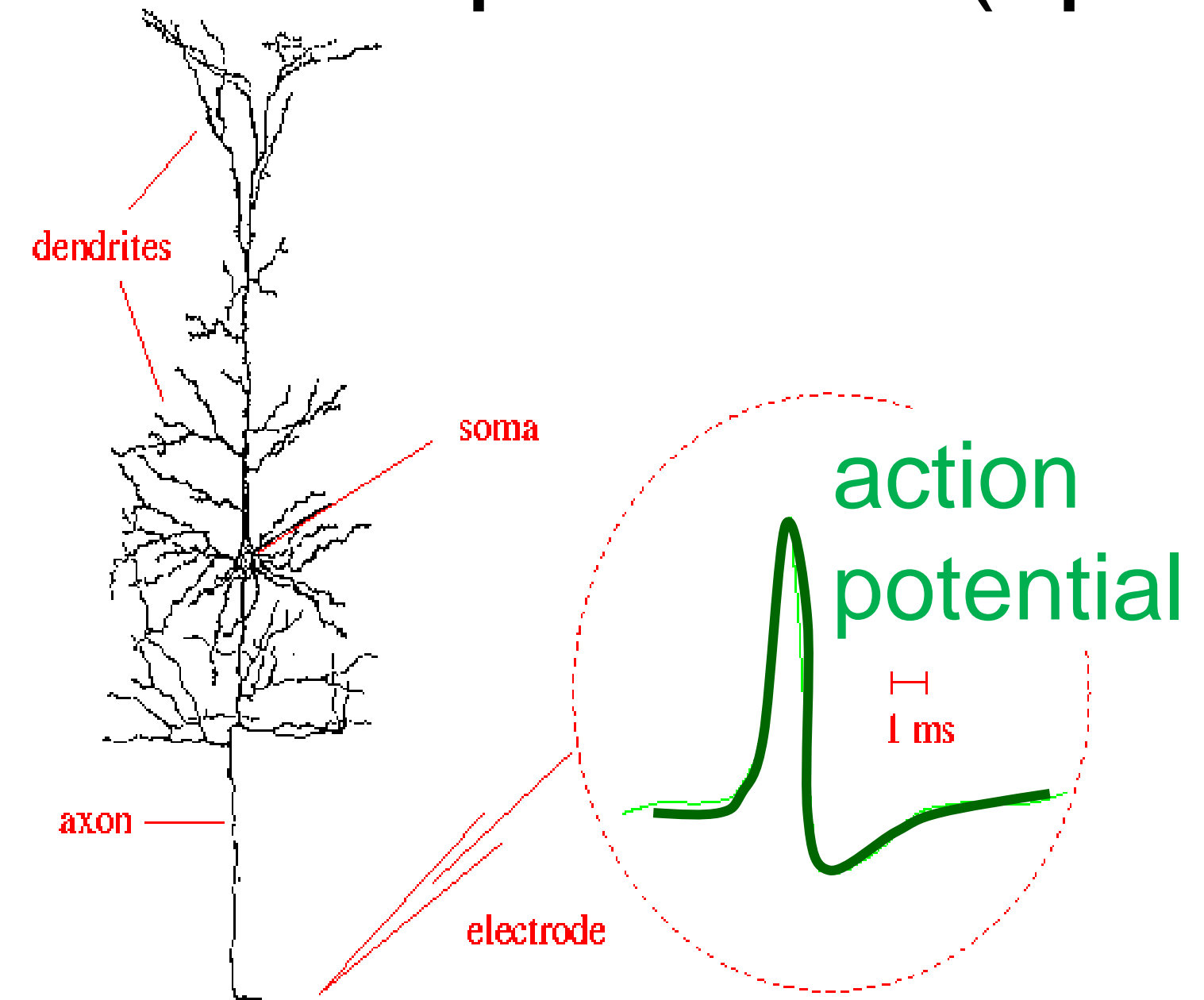
Hodgkin-Huxley type models:
Biophysics, molecules, ions
(week 2)

Signal:
action potential (spike)



Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

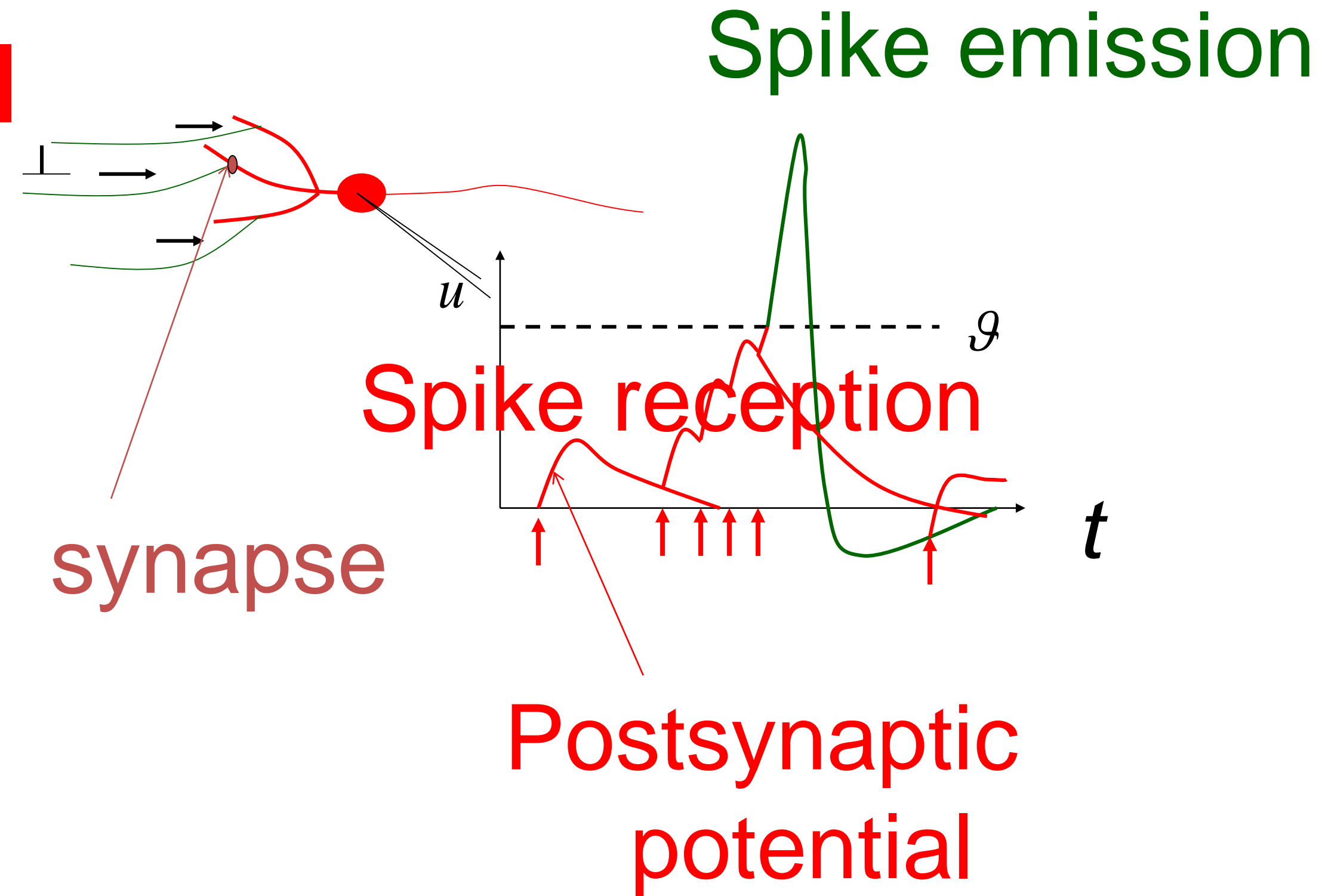
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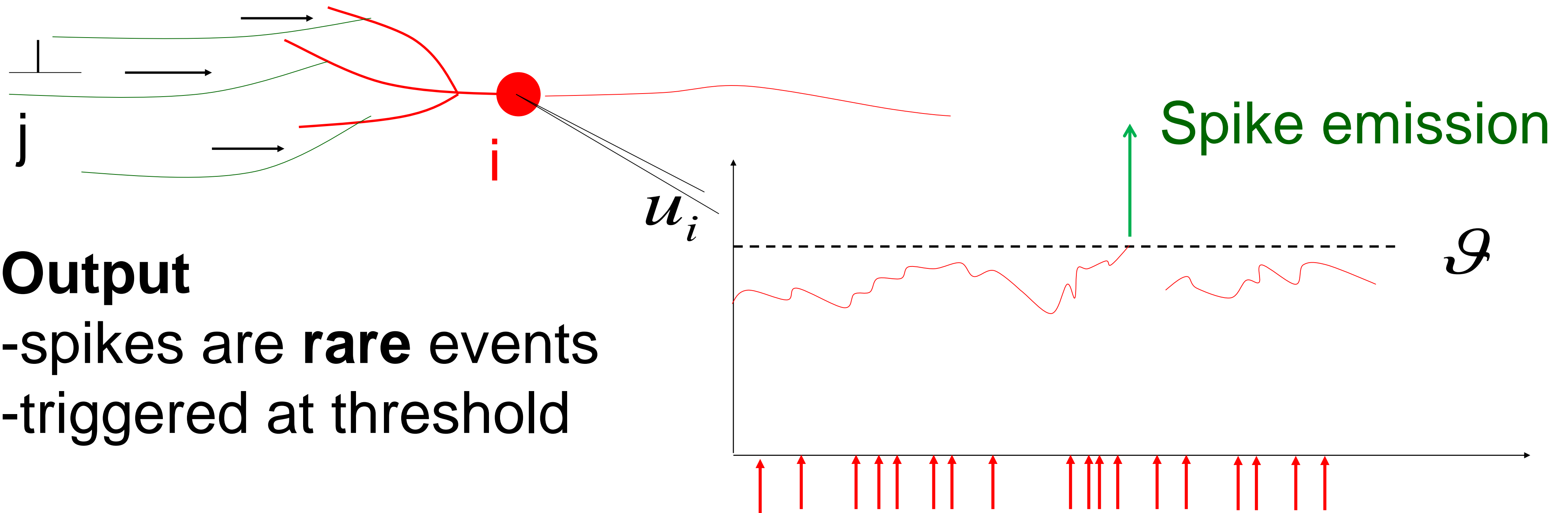
Neuronal Dynamics – 1.1. Neurons and Synapses/Overview

Integrate-and-fire models:
Formal/phenomenological
(week 1 and week 7-9)

- spikes are events
- triggered at threshold
- spike/reset/refractoriness



Noise and variability in integrate-and-fire models



Output

- spikes are **rare** events
- triggered at threshold

Subthreshold regime:

- trajectory of potential shows fluctuations

Random spike arrival

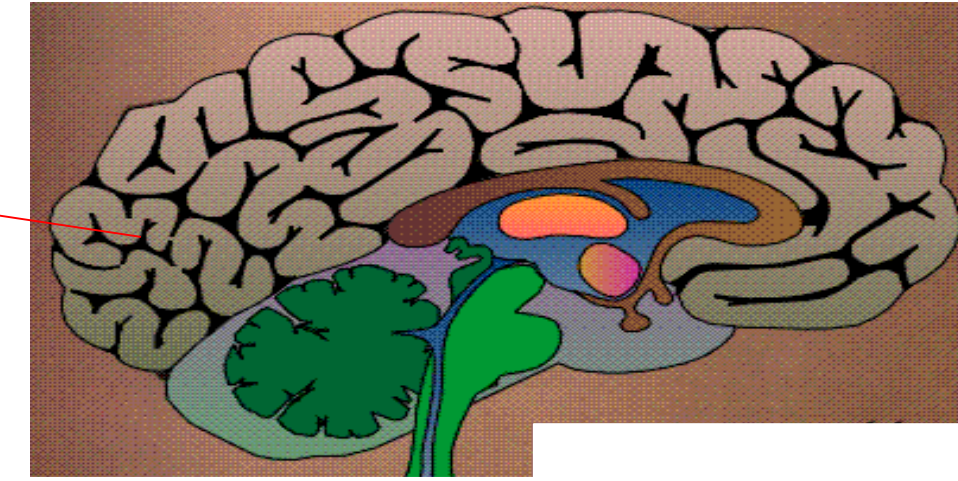
Neuronal Dynamics – membrane potential fluctuations

Spontaneous activity *in vivo*

What is noise?

What is the neural code?

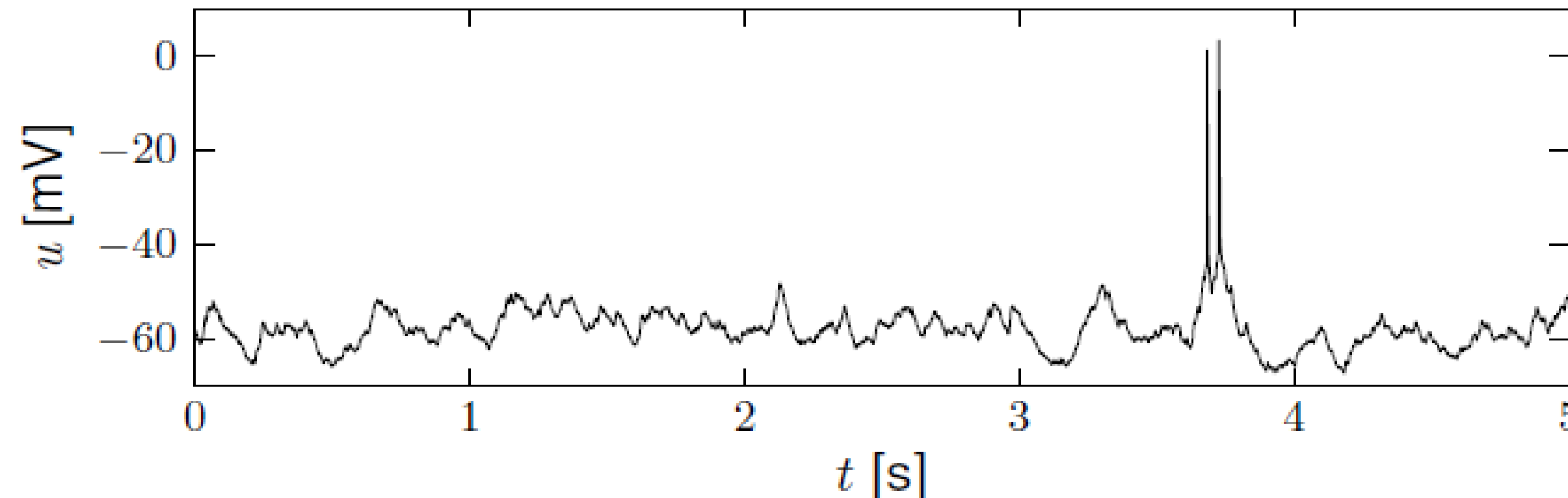
electrode



Brain

(week 11-13)

awake mouse, cortex, freely whisking,



Lab of Prof. C. Petersen, EPFL *Crochet et al., 2011*

Biological Modeling of Neural Networks – Quiz 1.1

A cortical neuron sends out signals which are called:

- action potentials
- spikes
- postsynaptic potential

The dendrite is a part of the neuron

- where synapses are located
- which collects signals from other neurons
- along which spikes are sent to other neurons

In an integrate-and-fire model, when the voltage hits the threshold:

- the neuron fires a spike
- the neuron can enter a state of refractoriness
- the voltage is reset
- the neuron explodes

In vivo, a typical cortical neuron exhibits

- rare output spikes
- regular firing activity
- a fluctuating membrane potential

Multiple answers possible!

Biological Modeling of Neural Networks

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EPFL, Lausanne, Switzerland

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biophysical modeling**

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Week 5,6: **Associative Memory,
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and coding**

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coding and decoding: GLM**

Week x: **Online video: Dendrites/Biophysics**

Biological modeling of Neural Networks

Course: Monday : 9:15-13:00

A typical Monday:

1st lecture 9:15-9:50

1st exercise 9:50-10:00

2nd lecture 10:15-10:35

2nd exercise 10:35-11:00

3rd lecture 11:15 – 11:40

3rd exercise 11:45-12:40

Course of 4 credits = 6 hours of work per week
4 'contact' + 2 homework

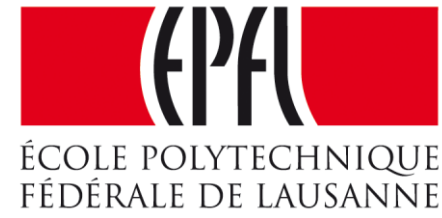
**have your laptop
with you**

paper and pencil

paper and pencil

*paper and pencil
OR interactive toy
examples
on computer*

Week 1 – part 2: The Passive Membrane



Biological Modeling of Neural Networks

**Week 1 – neurons and mathematics:
a first simple neuron model**

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EPFL, Lausanne, Switzerland

✓ 1.1 Neurons and Synapses:

Overview

1.2 The Passive Membrane

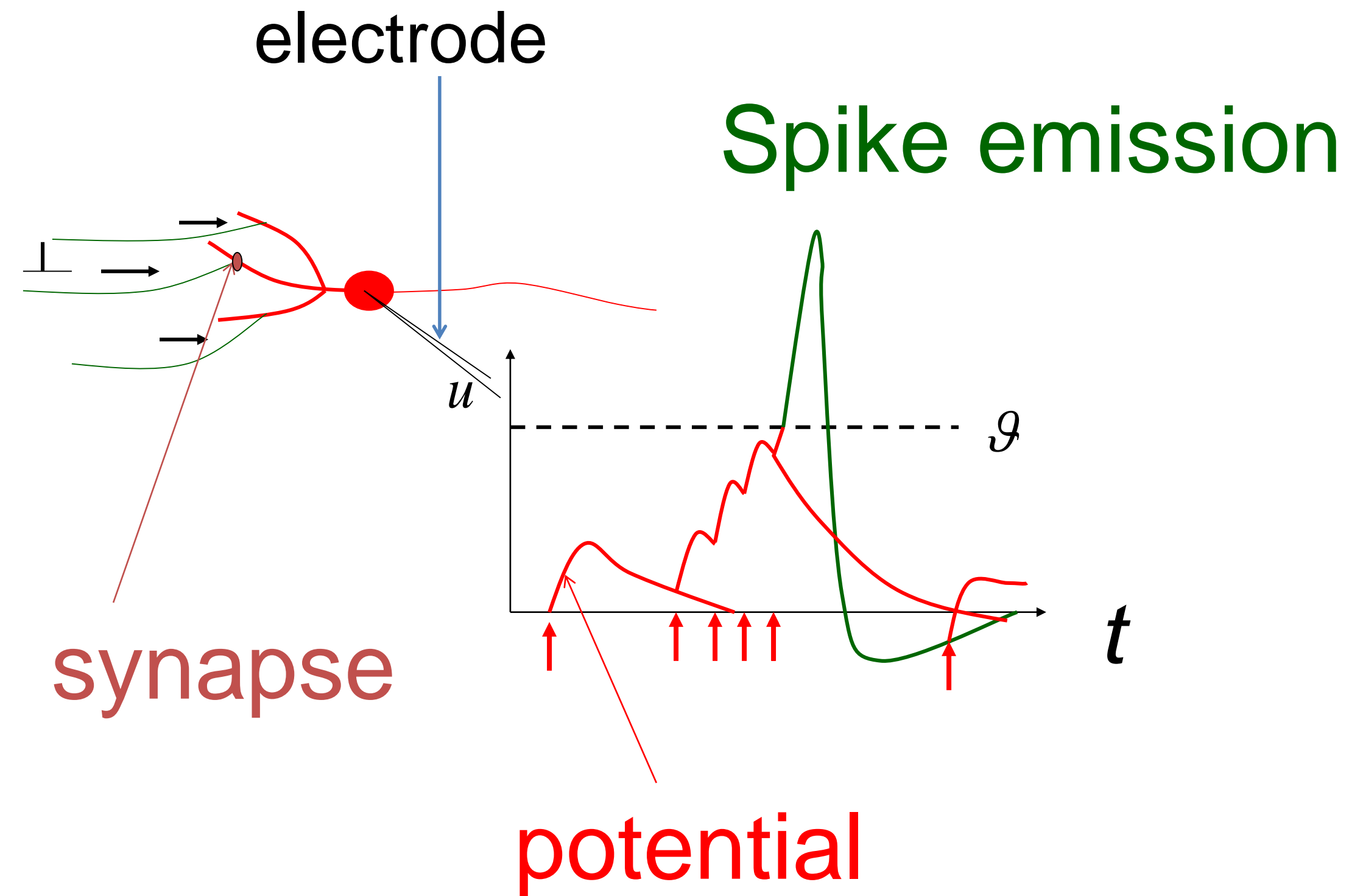
- Linear circuit
- Dirac delta-function

1.3 Leaky Integrate-and-Fire Model

1.4 Generalized Integrate-and-Fire Model

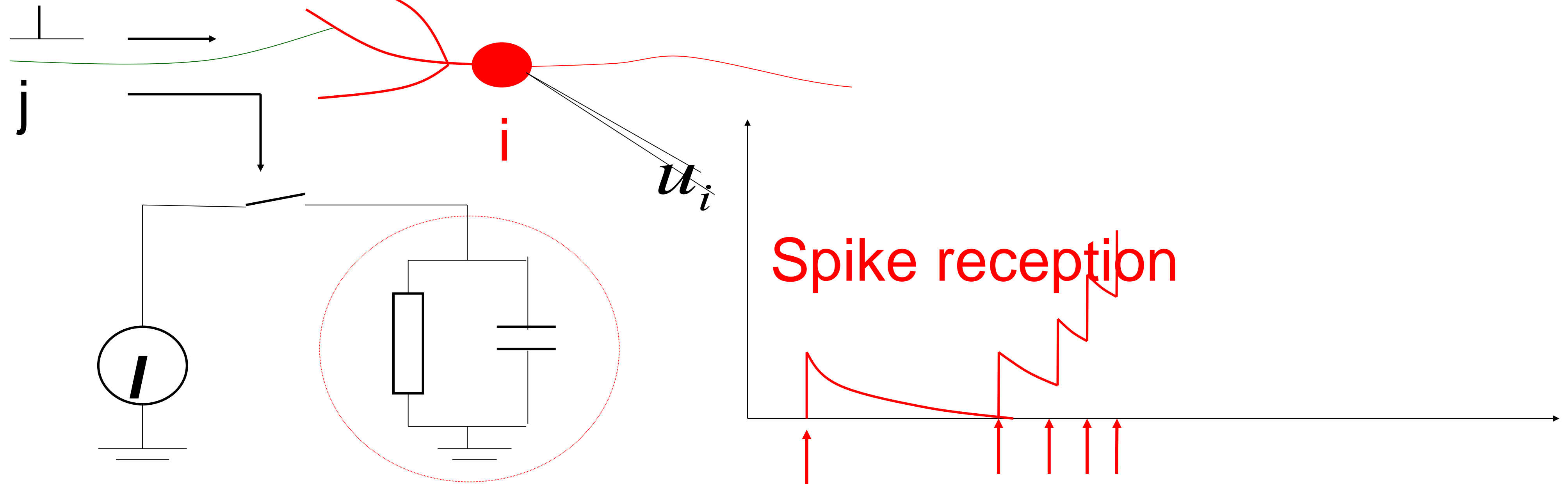
1.5. Quality of Integrate-and-Fire Models

Neuronal Dynamics – 1.2. The passive membrane



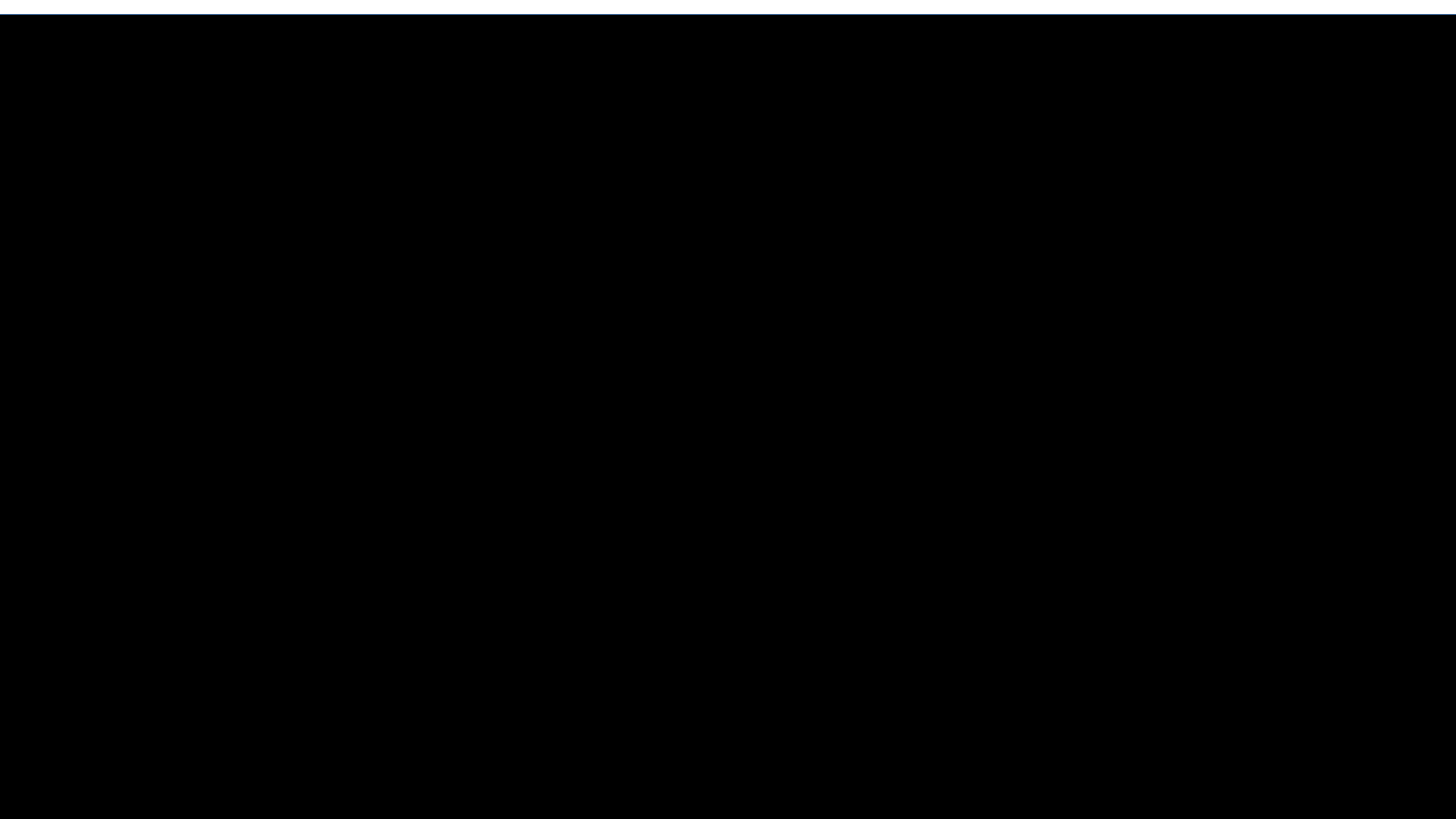
Integrate-and-fire model

Neuronal Dynamics – 1.2. The passive membrane

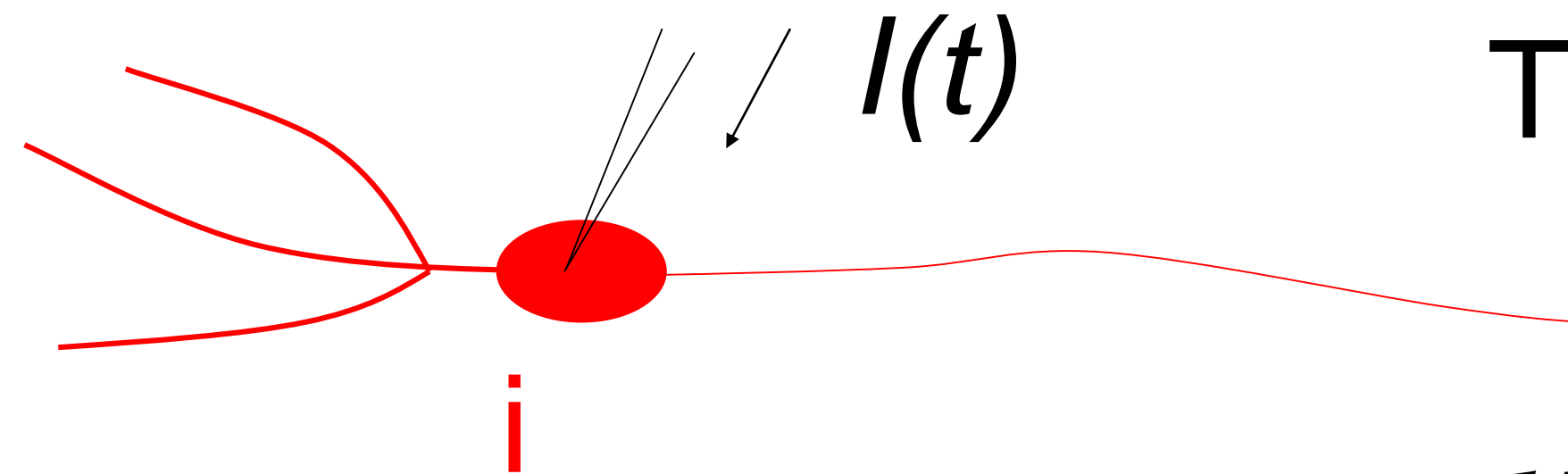


Subthreshold regime

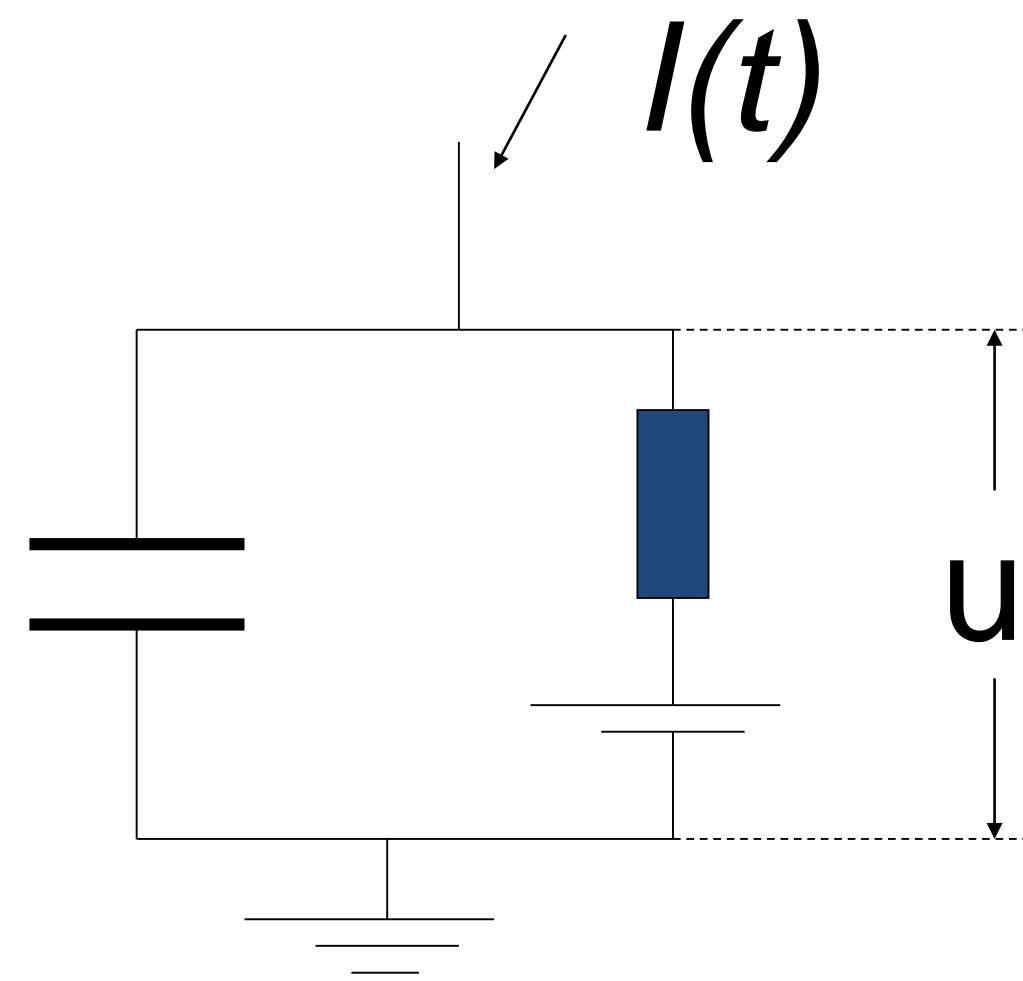
- linear
- passive membrane
- RC circuit



Neuronal Dynamics – 1.2. The passive membrane

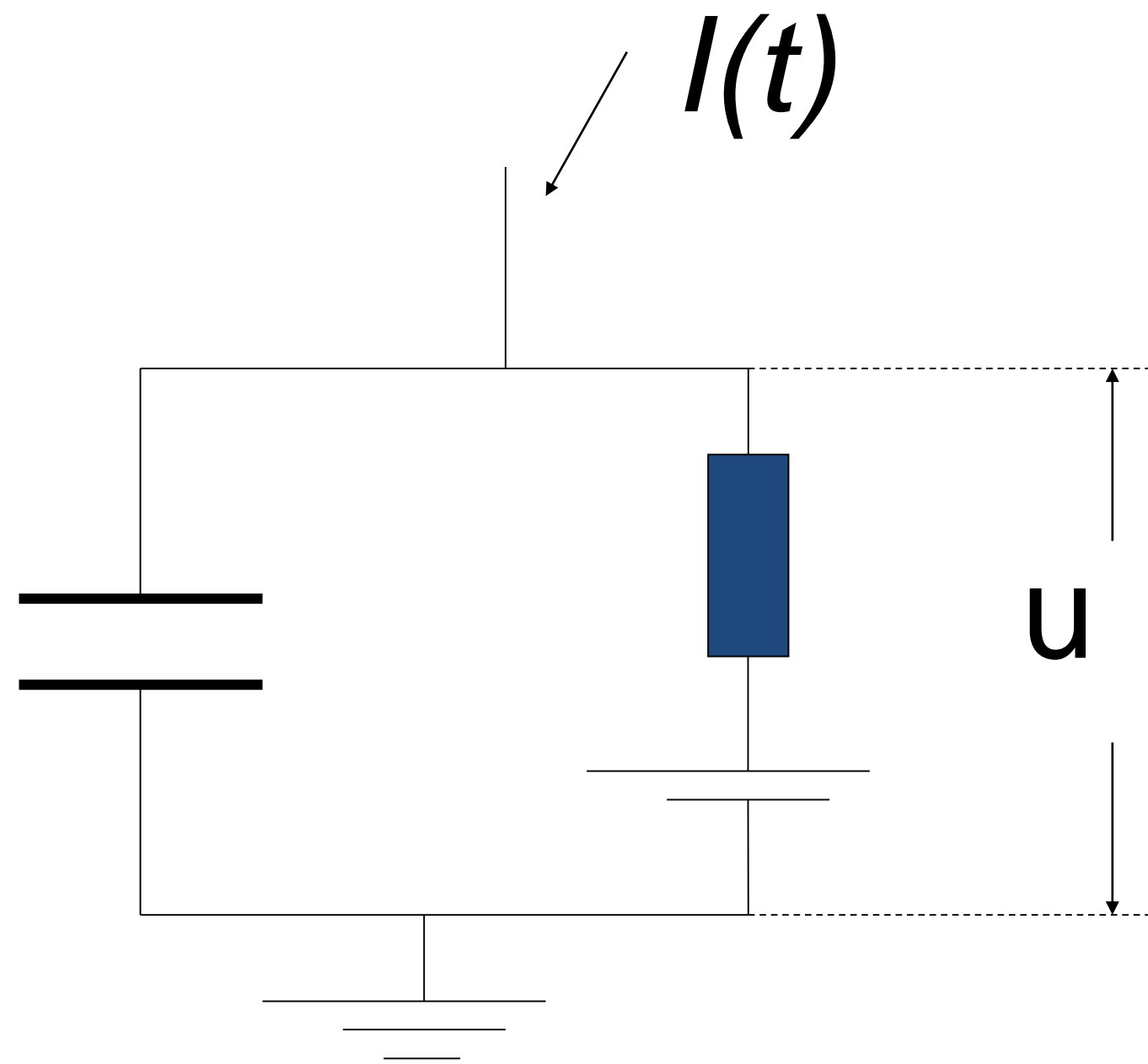


Time-dependent input

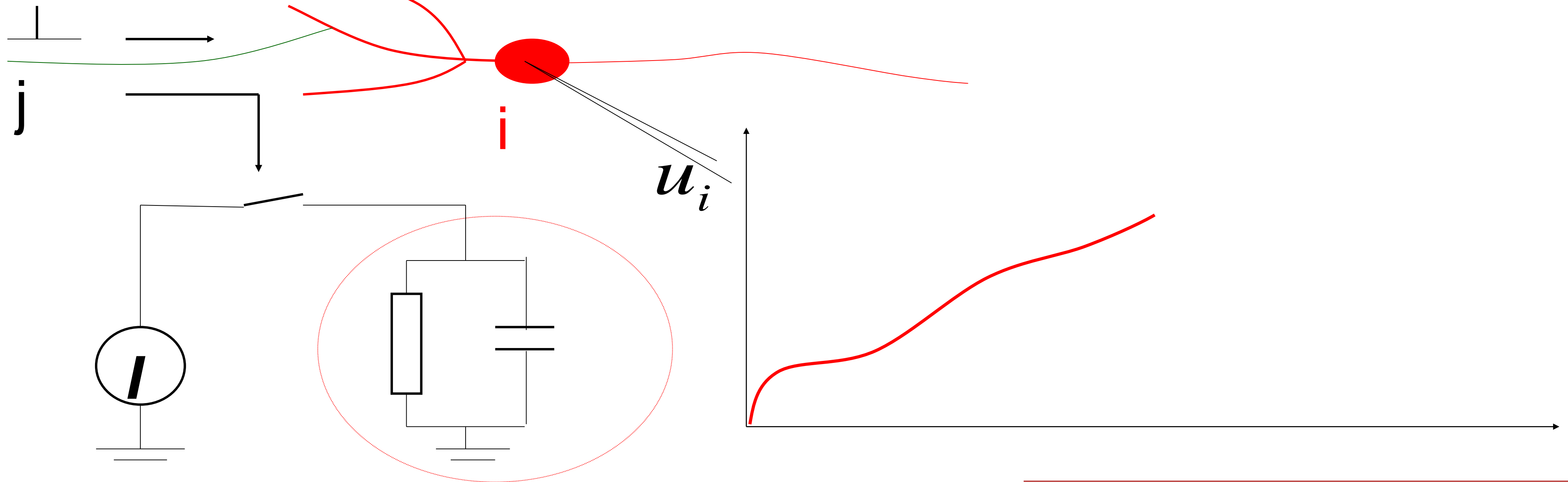


*Math development:
Derive equation
(Blackboard)*

Passive Membrane Model



Passive Membrane Model



$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

$$\tau \cdot \frac{d}{dt} V = -V + RI(t); \quad V = (u - u_{rest})$$

*Math Development:
Voltage rescaling
(blackboard)*

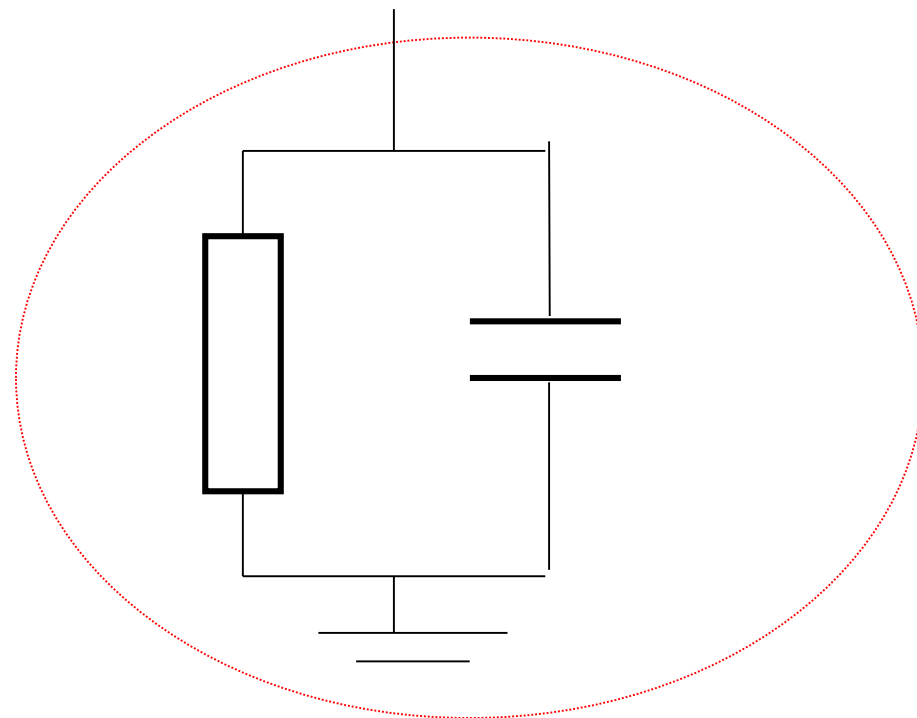
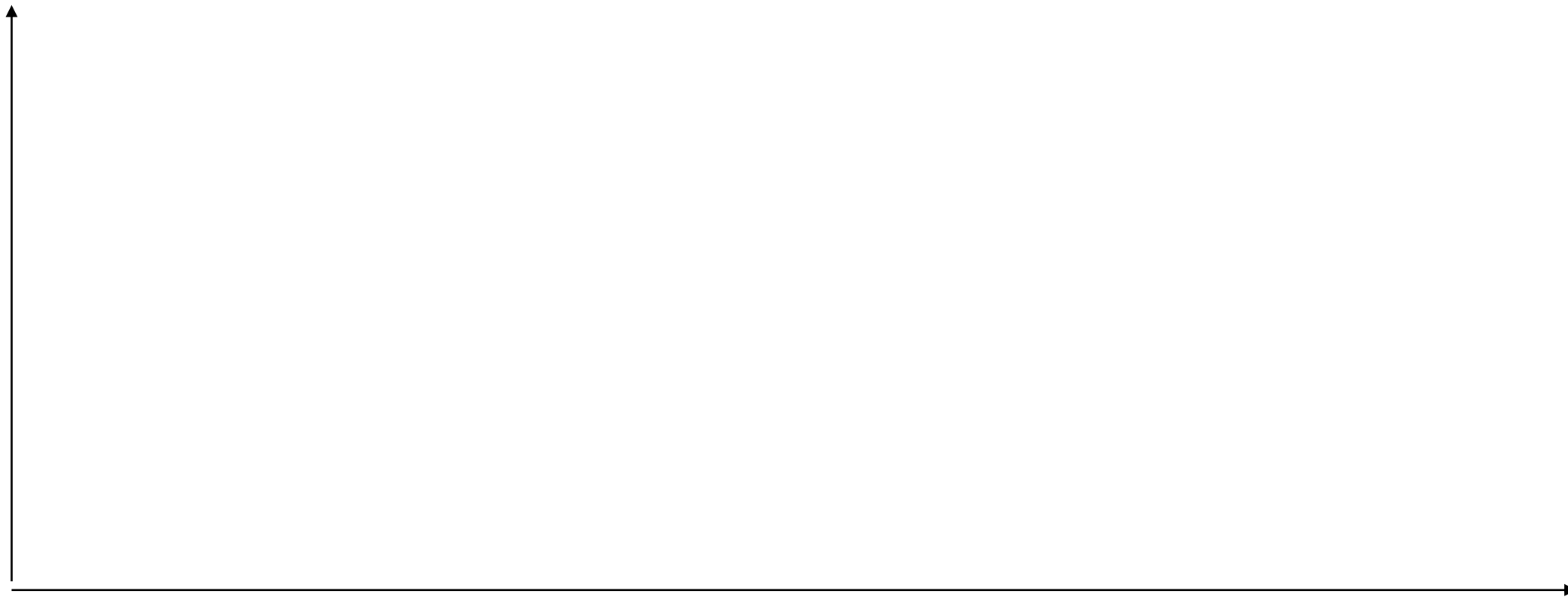
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Passive Membrane Model/Linear differential equation

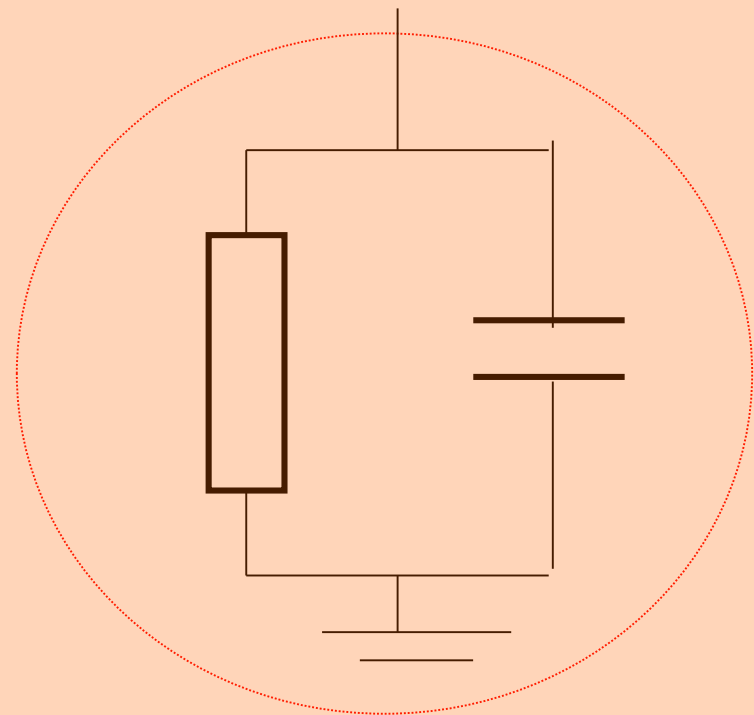
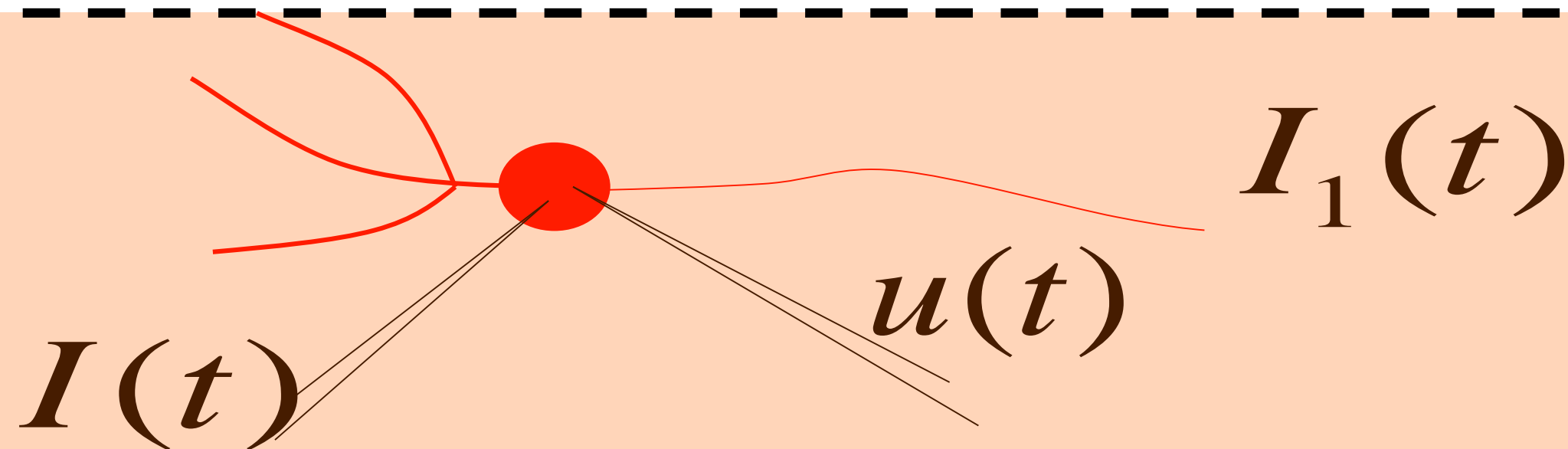
$$\tau \cdot \frac{d}{dt} V = -V + RI(t);$$



Free solution:
exponential decay

Neuronal Dynamics – Exercises NOW

Start Exerc. at 9:47.
Next lecture at
10:15



$I_2(t)$

$I_3(t)$

Step current input:

Pulse current input:

arbitrary current input:

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

$$\tau \cdot \frac{d}{dt} V = -V + RI(t); \quad V = (u - u_{rest})$$

**Calculate the voltage,
for the
3 input currents**

The voltage across a passive membrane can be described by the equation

$$\tau \frac{du}{dt} = -(u - u_{\text{rest}}) + RI(t). \quad (1)$$

Exercise 1: Passive Membrane

The voltage across a passive membrane can be described by the equation

$$\tau \frac{du}{dt} = -(u - u_{\text{rest}}) + RI(t). \quad (1)$$

1.1 Step current

Consider a current $I(t) = 0$ for $t < t_0$ and $I(t) = I_0$ for $t > t_0$. Calculate the voltage $u(t)$, given that the neuron is at rest at time t_0 . (Hint: Instead of solving the differential equation explicitly, try to construct the response to the step current along the lines: What is the value of $u(t)$ for $t \leq t_0$? What is the asymptotic value of $u(t)$ for $t \gg t_0$? What is the functional form and time scale of the transition?)

1.2 Pulse current

Consider a current pulse

$$I(t) = \begin{cases} 0 & \text{for } t < t_0 \text{ and } t > t_0 + \Delta \\ q/\Delta & \text{for } t \geq t_0 \text{ and } t < t_0 + \Delta, \end{cases} \quad (2)$$

where Δ is a short time and q is the total electrical charge.

Consider first $\Delta = 0.1\tau$, and then $\Delta = 0.05\tau$, $\Delta = 0.025\tau$. Sketch the input current pulse and the voltage response. What happens in the limit $\Delta \rightarrow 0$? (Hint: Use $e^{-x} \approx 1 - x$ for $x \ll 1$.)

1.3 Delta function

The Dirac delta function can be defined by the limit of a short pulse:

$$\delta(t - t_0) = \lim_{\Delta \rightarrow 0} f_{\Delta}(t) \quad \text{where} \quad f_{\Delta}(t) = \begin{cases} 1/\Delta & \text{for } t_0 \leq t < t_0 + \Delta \\ 0 & \text{otherwise.} \end{cases} \quad (3)$$

Convince yourself that the integral $\int_{t_1}^{t_2} \delta(t - t_0) dt$ is equal to one if $t_1 \leq t_0 < t_2$ and vanishes otherwise.

Express $I(t)$ in Eq. 1 using the δ -function for the case that an extremely short current pulse arrives at time t^f . Pay attention to the units!

1.4 General solution

Assuming that before a given time t_0 the current is null and the membrane potential is at rest, derive the general solution to Eq. (1) for arbitrary $I(t)$.

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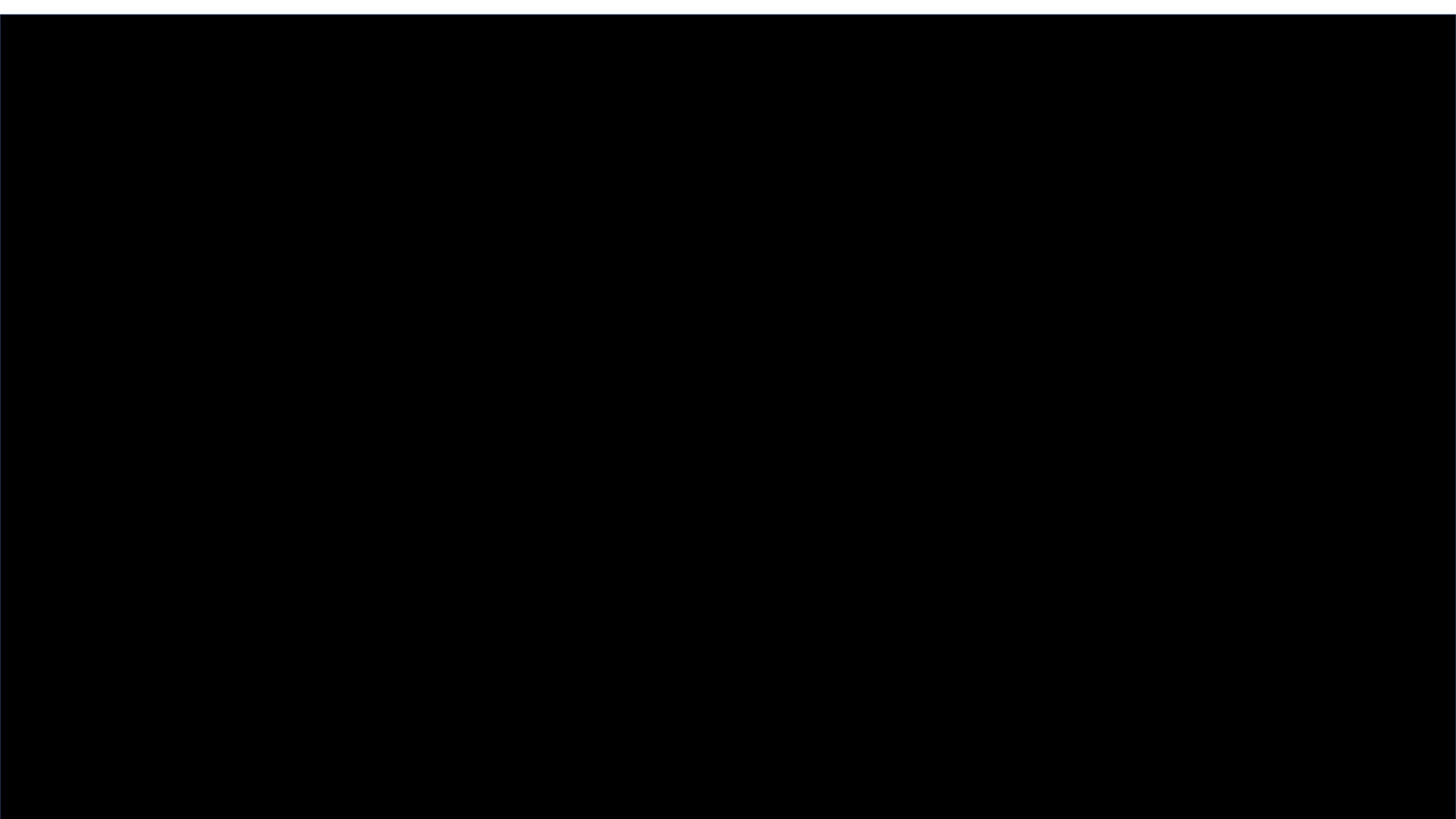
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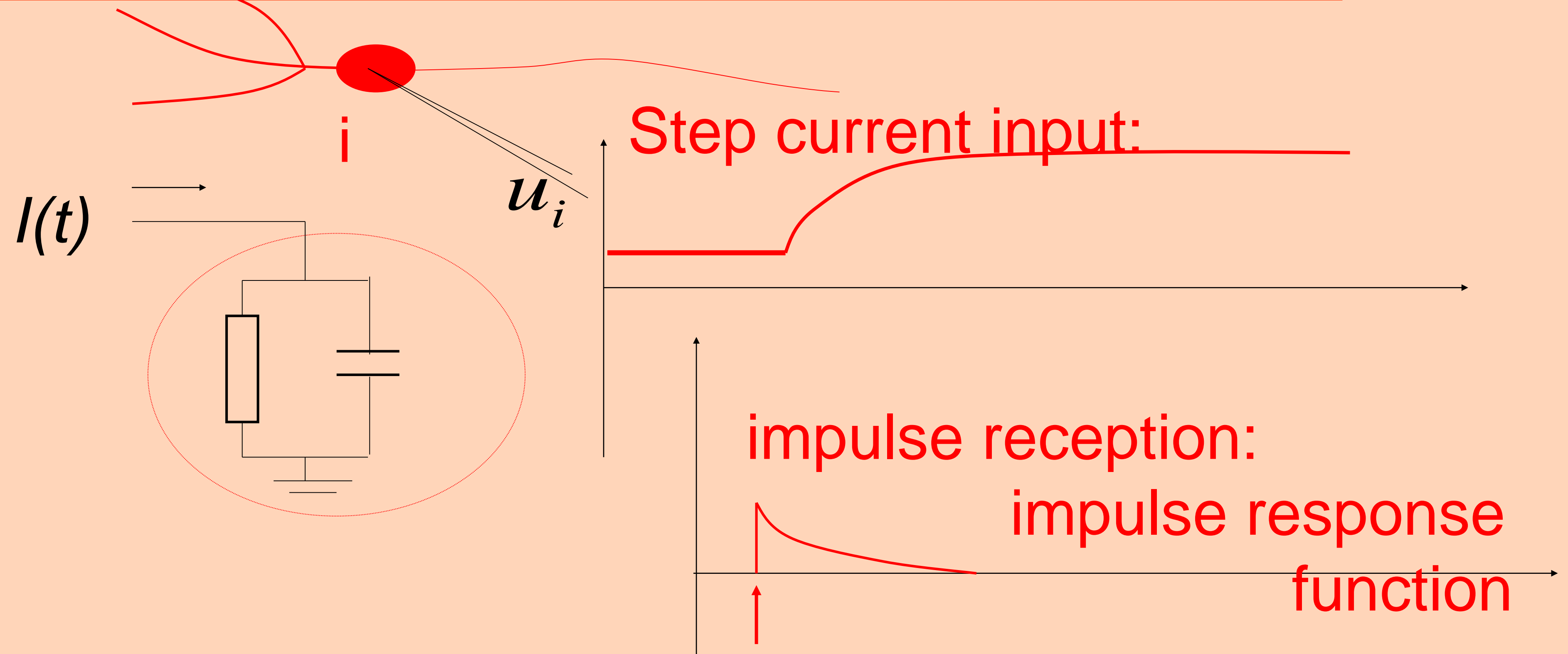
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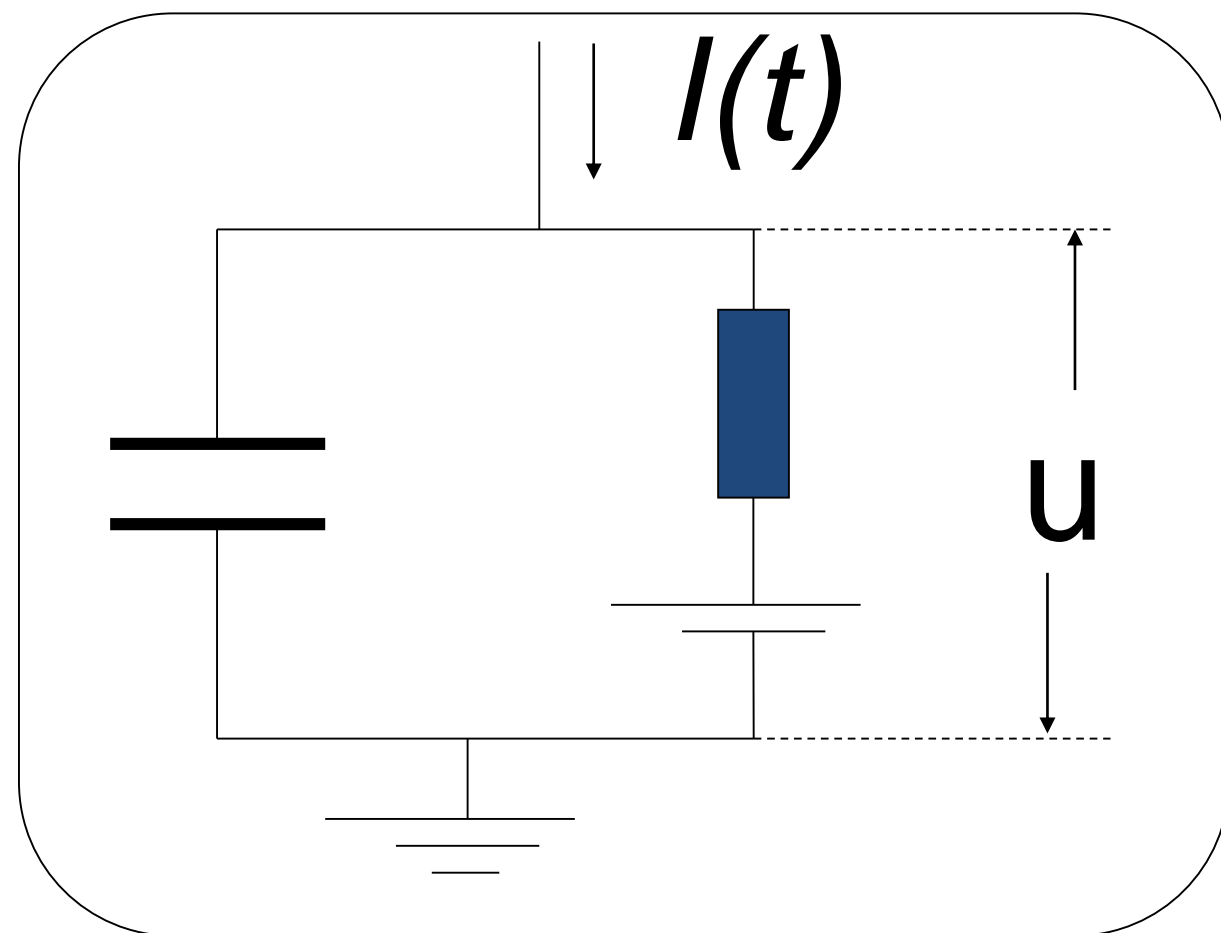
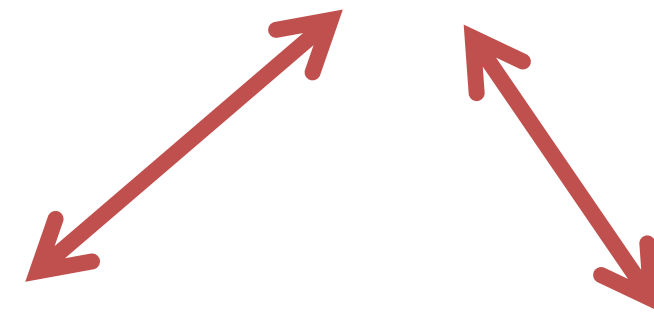
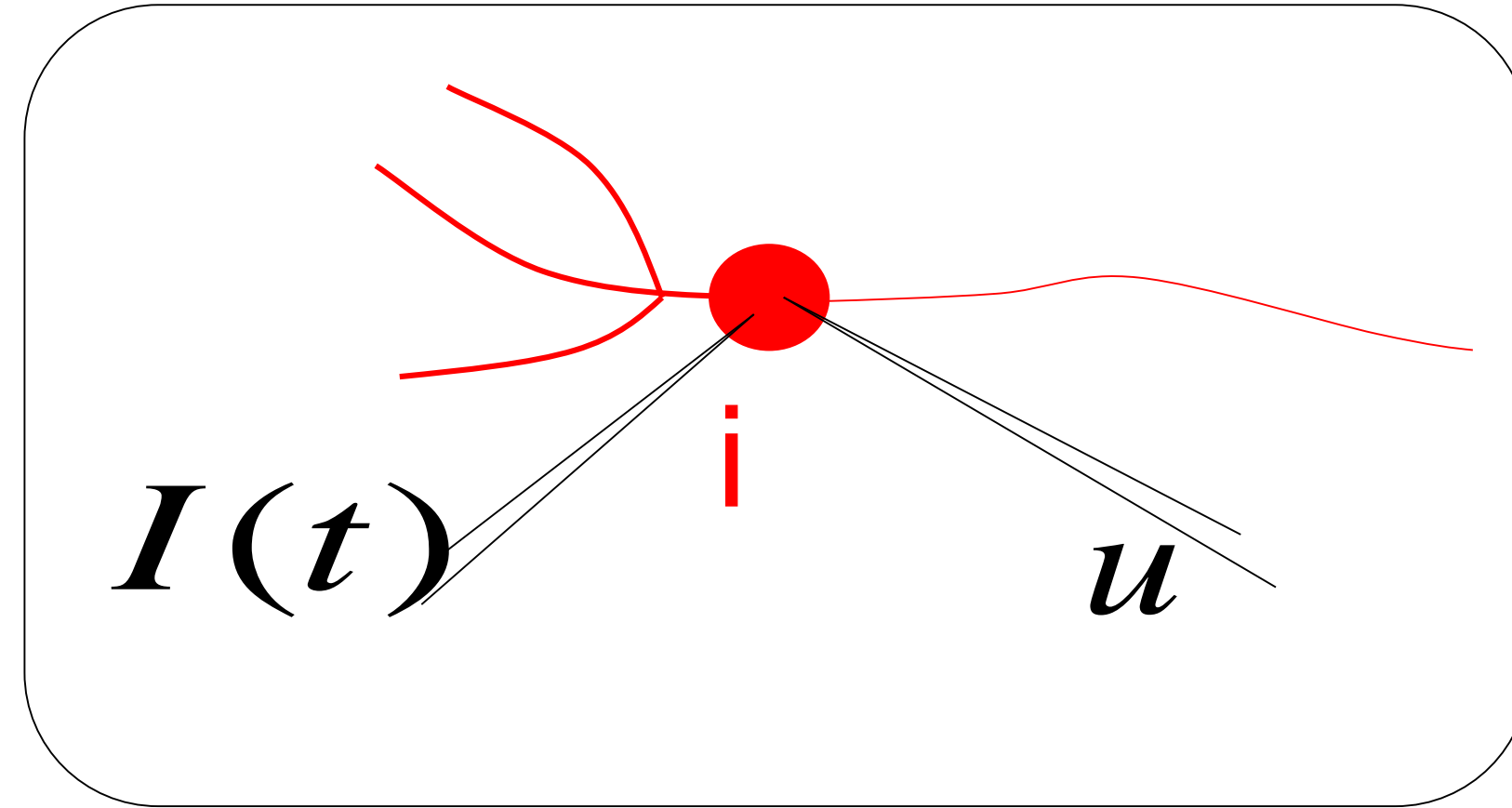
Passive Membrane Model – exercise 1 now



Linear equation

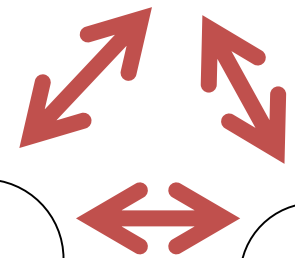
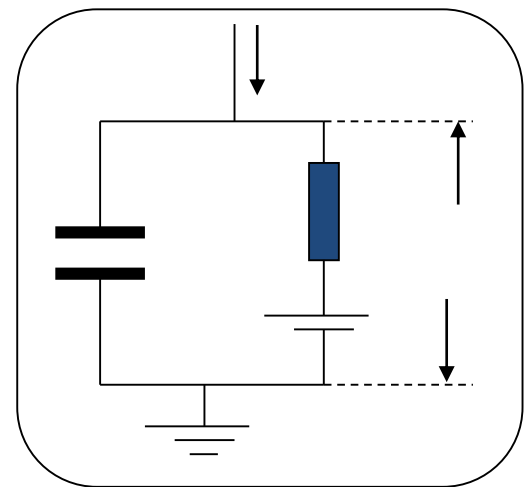
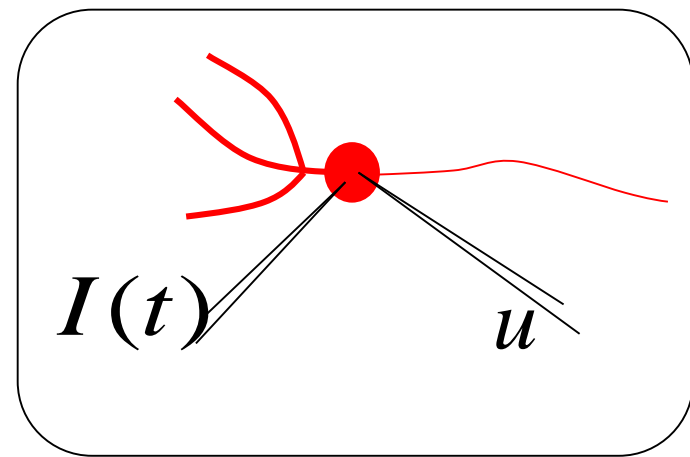
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Triangle: neuron – electricity - math

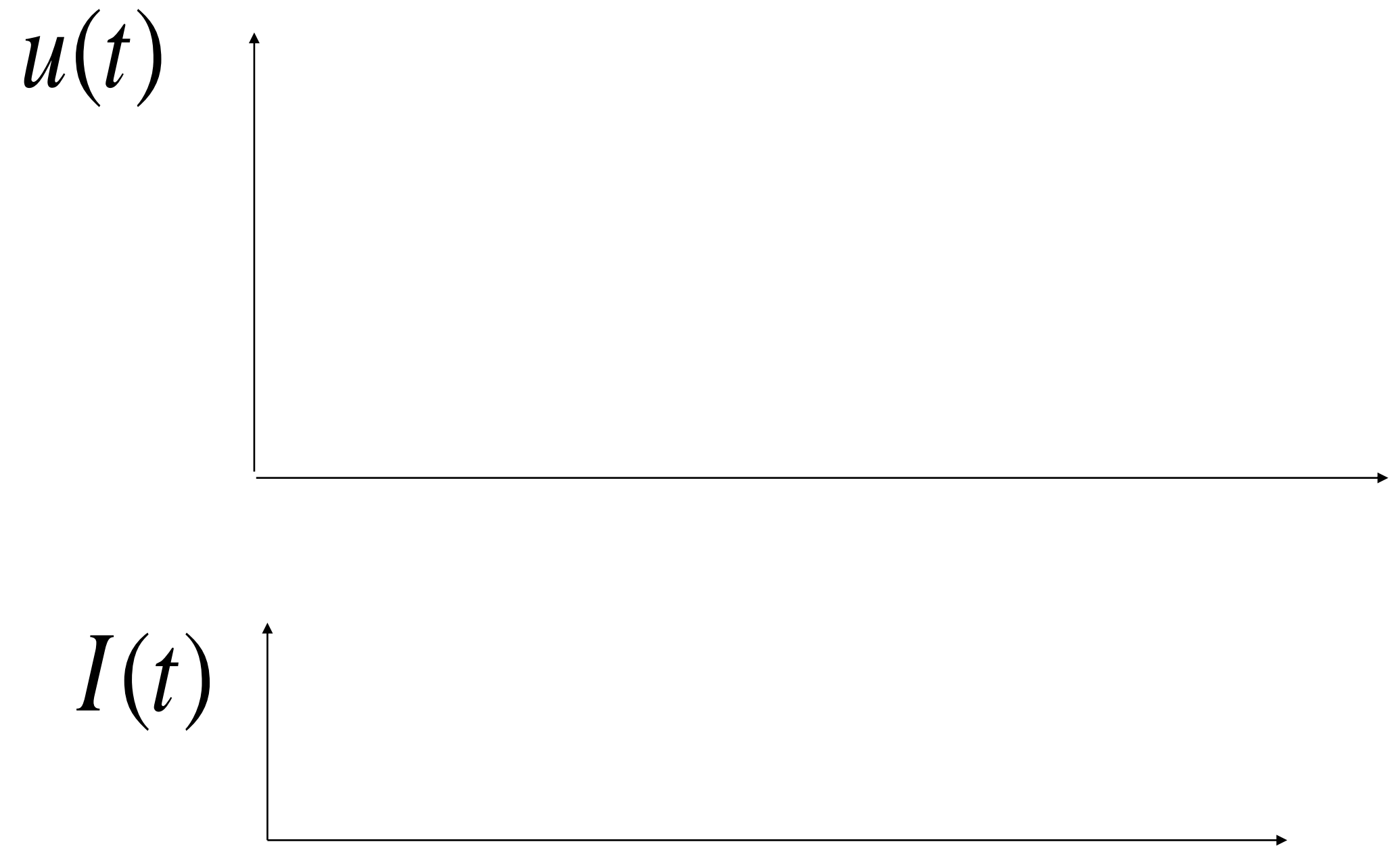


$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Pulse input – charge – delta-function



$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

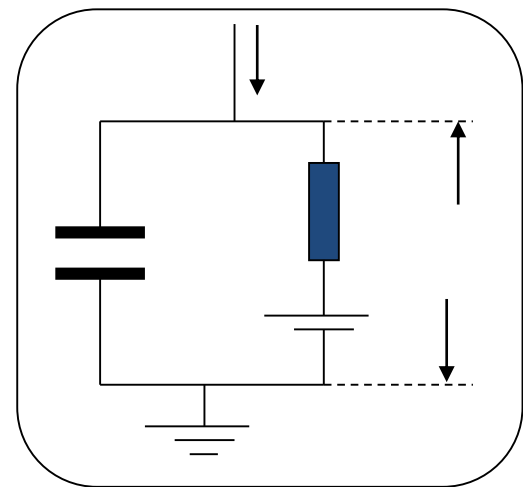
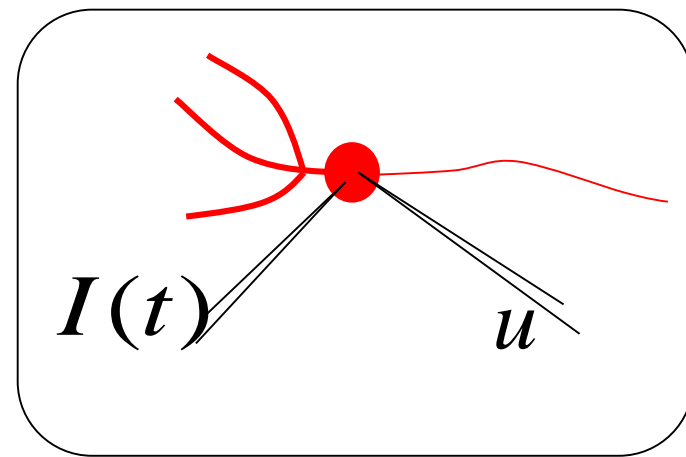


$$I(t) = q \cdot \delta(t - t_0)$$

Pulse current input

Dirac delta-function

$$I(t) = q \cdot \delta(t - t_0)$$



$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

$$1 = \int_{t_0-a}^{t_0+a} \delta(t - t_0) dt$$
$$f(t_0) = \int_{t_0-a}^{t_0+a} f(t) \delta(t - t_0) dt$$

Neuronal Dynamics – Solution of Ex. 1 – arbitrary input

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Arbitrary input

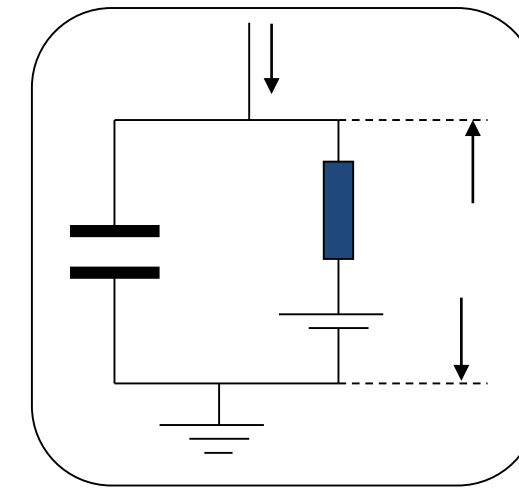
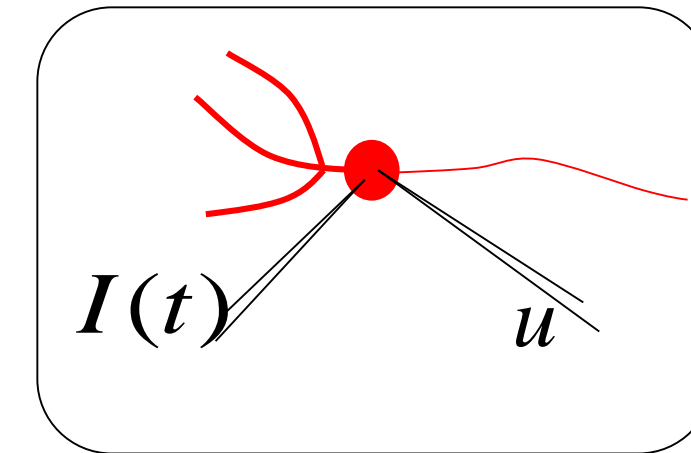
$$u(t) = u_{rest} + \int_{-\infty}^t \frac{1}{c} e^{-(t-t')/\tau} I(t') dt'$$

Single pulse

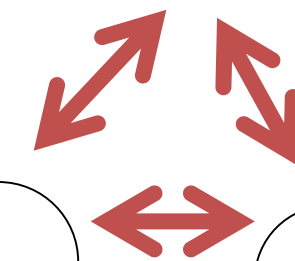
$$\Delta u(t) = \frac{q}{c} e^{-(t-t_0)/\tau}$$

you need to know the solutions of linear differential equations!

Passive membrane, linear differential equation

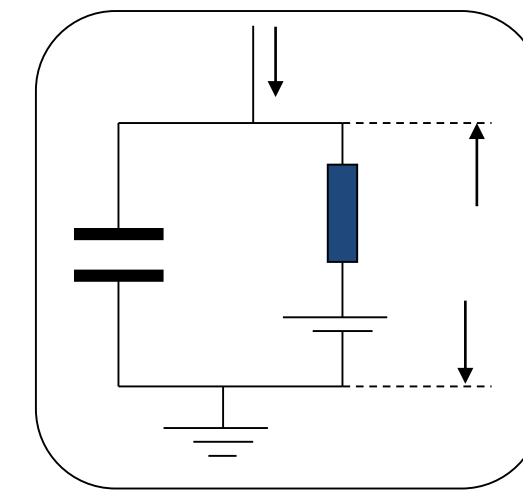
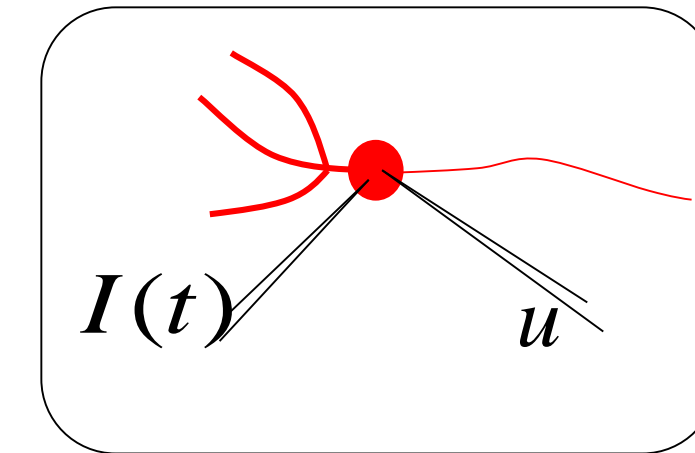


$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$



Passive membrane, linear differential equation

*If you have difficulties,
watch lecture 1.2detour.*



$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Three prerequisites:

- Analysis 1-3
- Probability/Statistics
- Differential Equations or Physics 1-3 or Electrical Circuits

<https://lcnwww.epfl.ch/gerstner/NeuronalDynamics-MOOC1.html>

LEARNING OUTCOMES

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Transversal skills

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Use a textbook,
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miniproject

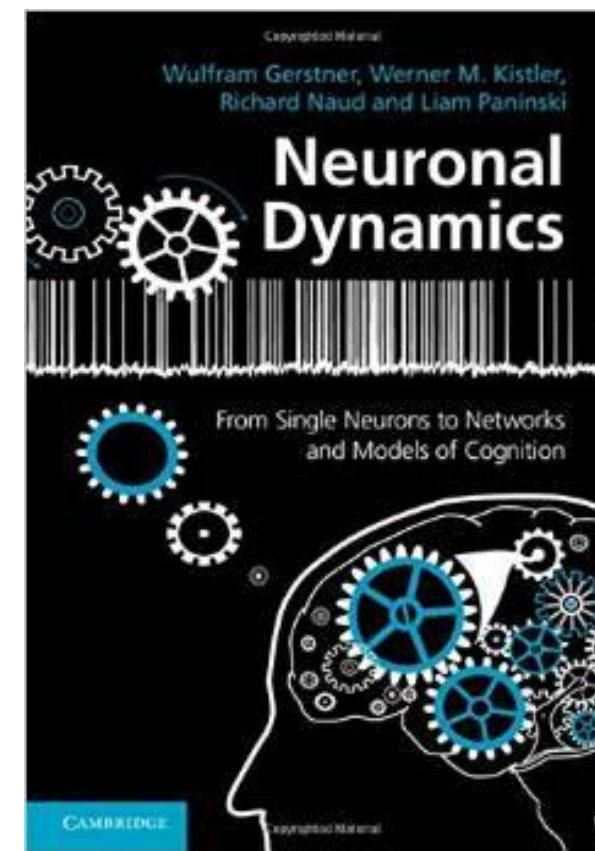
Biological Modeling of Neural Networks

Written Exam (70%)
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of which you have to hand in 2

Textbook:

<http://neurondynamics.epfl.ch/>



Video:

<https://lcnwww.epfl.ch/gerstner/NeuronalDynamics-MOOC1.html>

<https://lcnwww.epfl.ch/gerstner/NeuronalDynamics-MOOC2.html>

Biological Modeling of Neural Networks

Questions?

This is **not** a course on

Deep learning or **Artificial neural networks**

→ Deep Learning, master EE, (*Fleuret*)

→ Artificial NN, master CS, (*Gerstner*)

Summary of Section 1.1 and 1.2.

Neurons emit spikes (action potentials) which are short standardized events in the form of voltage pulses.

Spikes are emitted at the firing threshold.

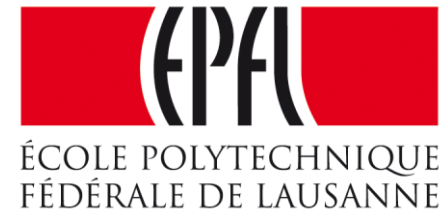
Below the threshold the electrical behavior is often well characterized by a linear differential equation (math)

corresponding to an RC circuit (electricity) or to a passive membrane (biology). We will often in this class walk along the triangle that connects math with electricity and biology.

This class has a strong focus on mathematical modeling of the biological phenomena. Differential equations, Dirac-delta pulses, and their link to biology are important concepts.

The time constant of an RC circuit is $\tau=RC$.

Week 1 – part 3: Leaky Integrate-and-Fire Model



Biological Modeling of Neural Networks

**Week 1 – neurons and mathematics:
a first simple neuron model**

Wulfram Gerstner

EPFL, Lausanne, Switzerland

√ 1.1 Neurons and Synapses:
Overview

√ 1.2 The Passive Membrane

- Linear circuit
- Dirac delta-function
- Detour: solution of 1-dim linear differential equation

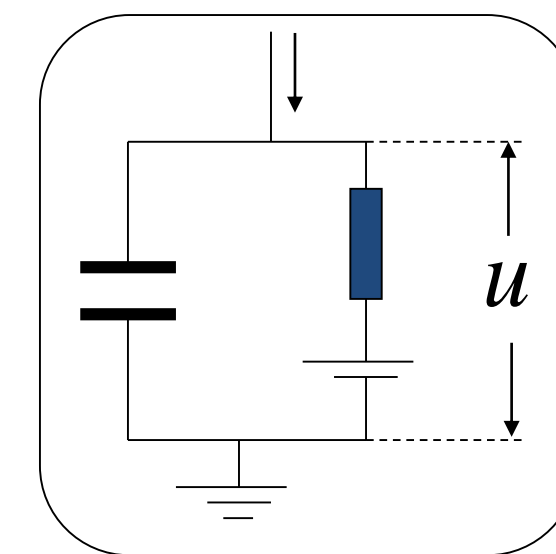
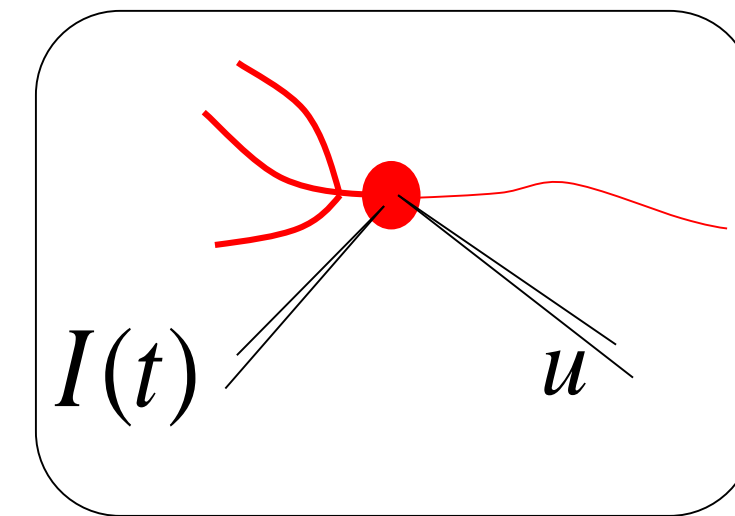
1.3 Leaky Integrate-and-Fire Model

1.4 Generalized Integrate-and-Fire Model

1.5. Quality of Integrate-and-Fire Models

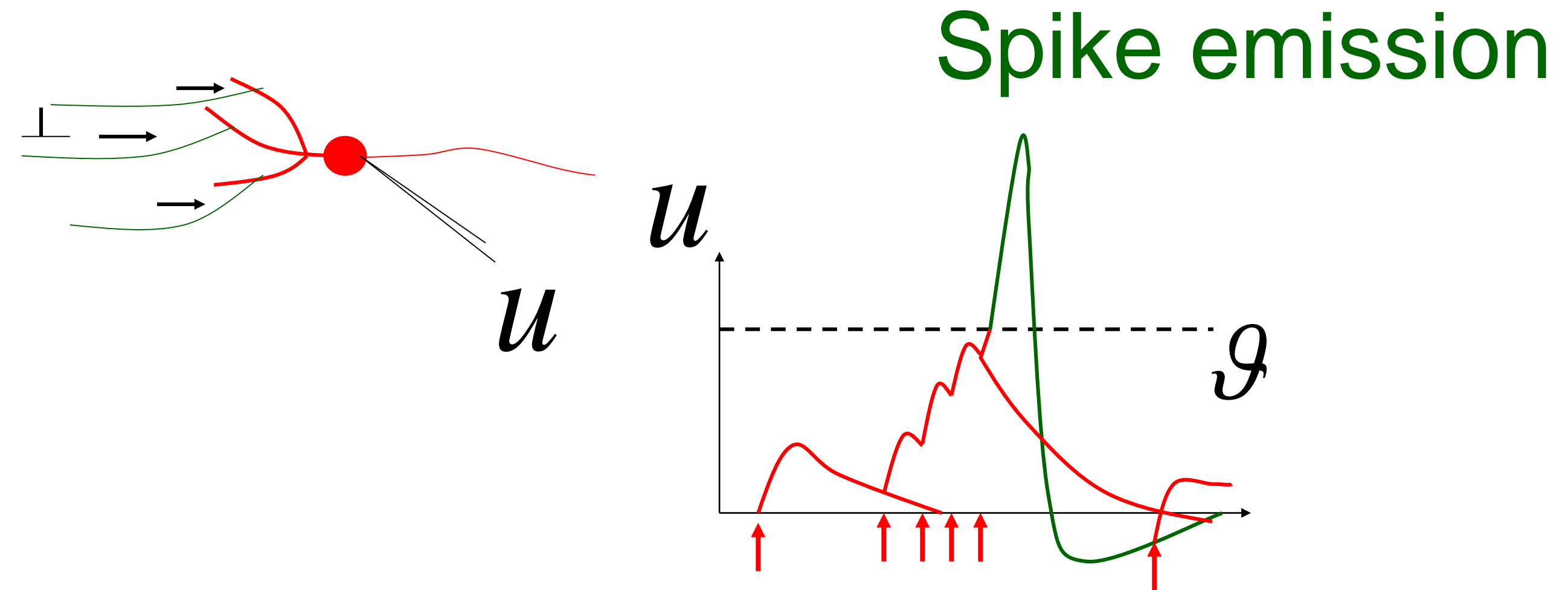
Neuronal Dynamics – 1.3 Leaky Integrate-and-Fire Model

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$



$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Neuronal Dynamics – Integrate-and-Fire type Models



Simple

Integrate-and-Fire Model:

passive membrane
+ *threshold*

Input spike causes an EPSP
= excitatory postsynaptic potential

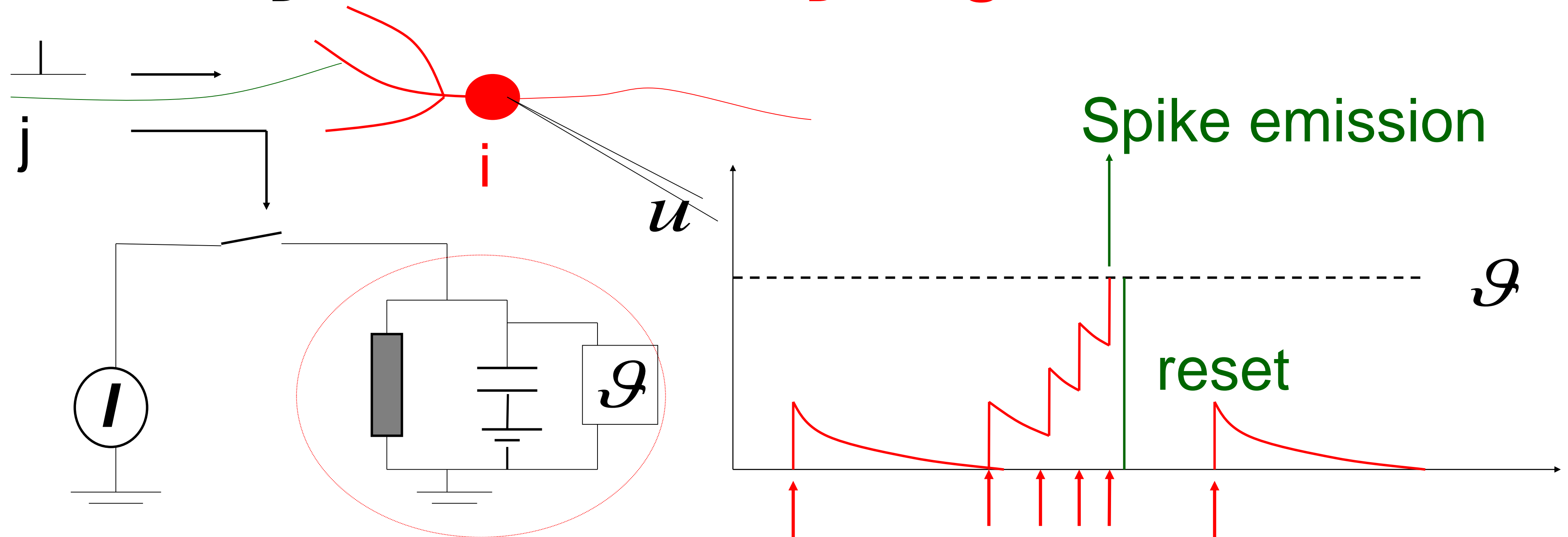
-output spikes are events

-generated at threshold

-after spike: reset/refractoriness

Leaky Integrate-and-Fire Model

Neuronal Dynamics – 1.3 Leaky Integrate-and-Fire Model



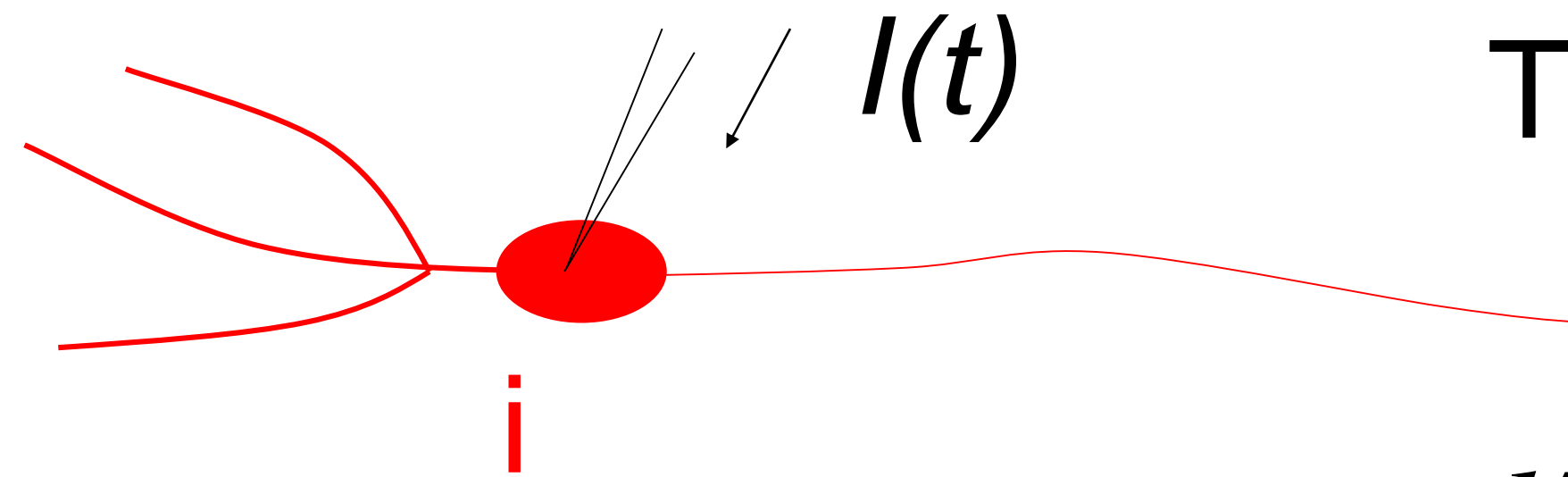
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

linear

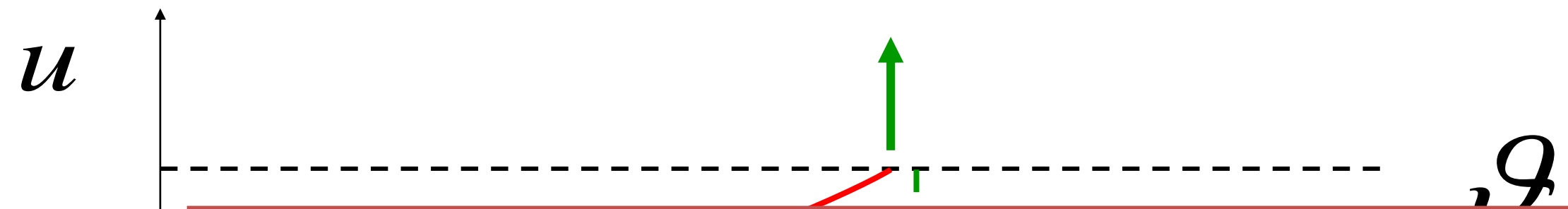
$$u(t) = \mathcal{G} \Rightarrow \text{Fire+reset } u \rightarrow u_r$$

threshold

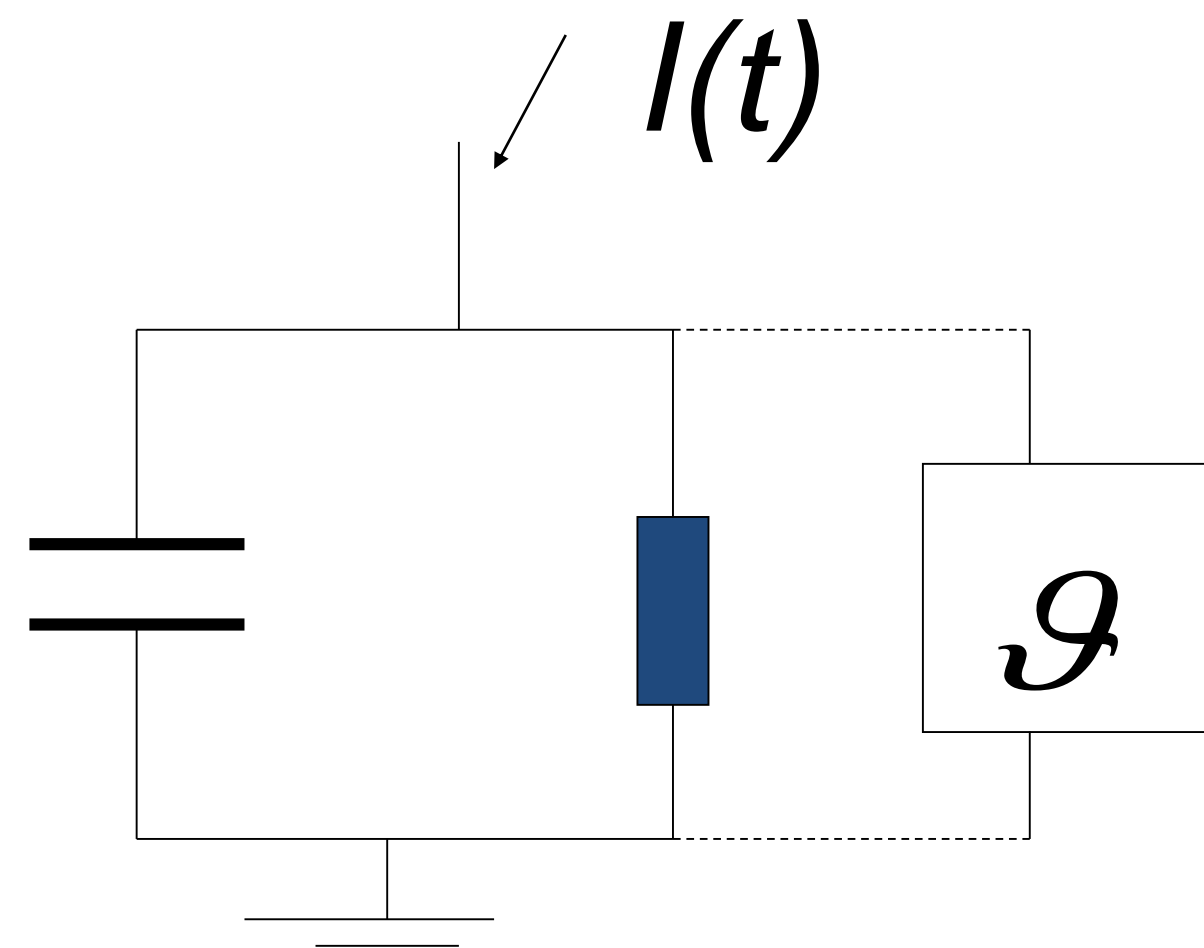
Neuronal Dynamics – 1.3 Leaky Integrate-and-Fire Model



Time-dependent input



*Math development:
Response to step current*



- spikes are events
- triggered at threshold
- spike/reset/refractoriness

Week1 – Quiz 2.

Take 90 seconds:

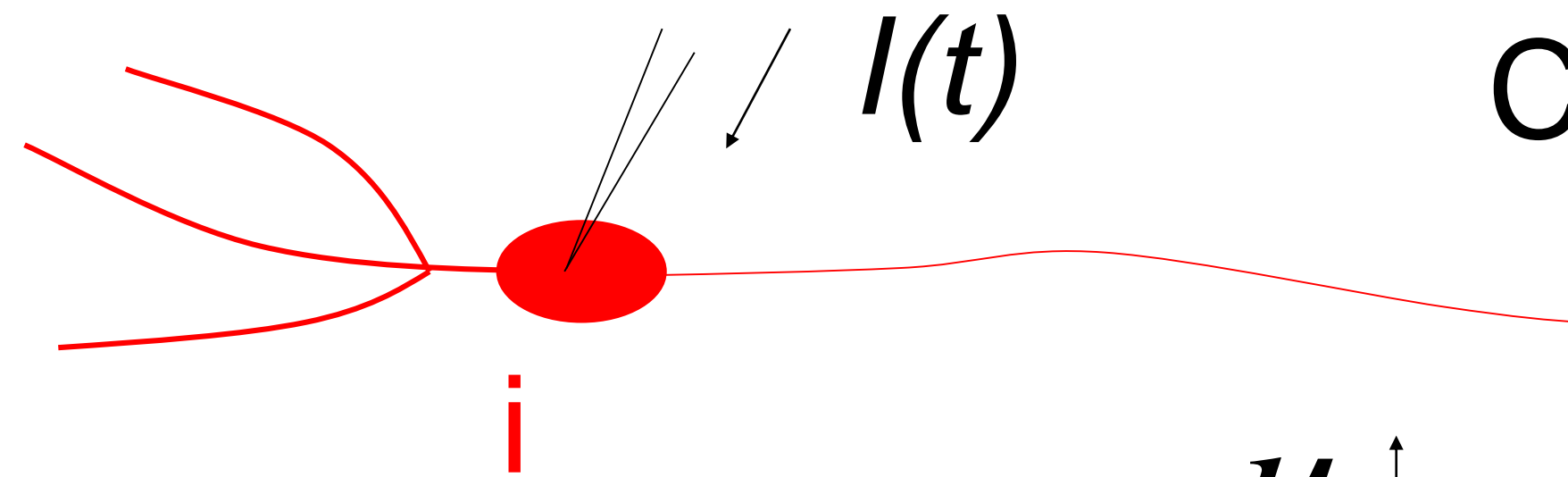
Consider the linear differential equation $\tau \cdot \frac{d}{dt} x = -x + x_c$
with initial condition at $t = 0 : x = 0$

The solution for $t > 0$ is

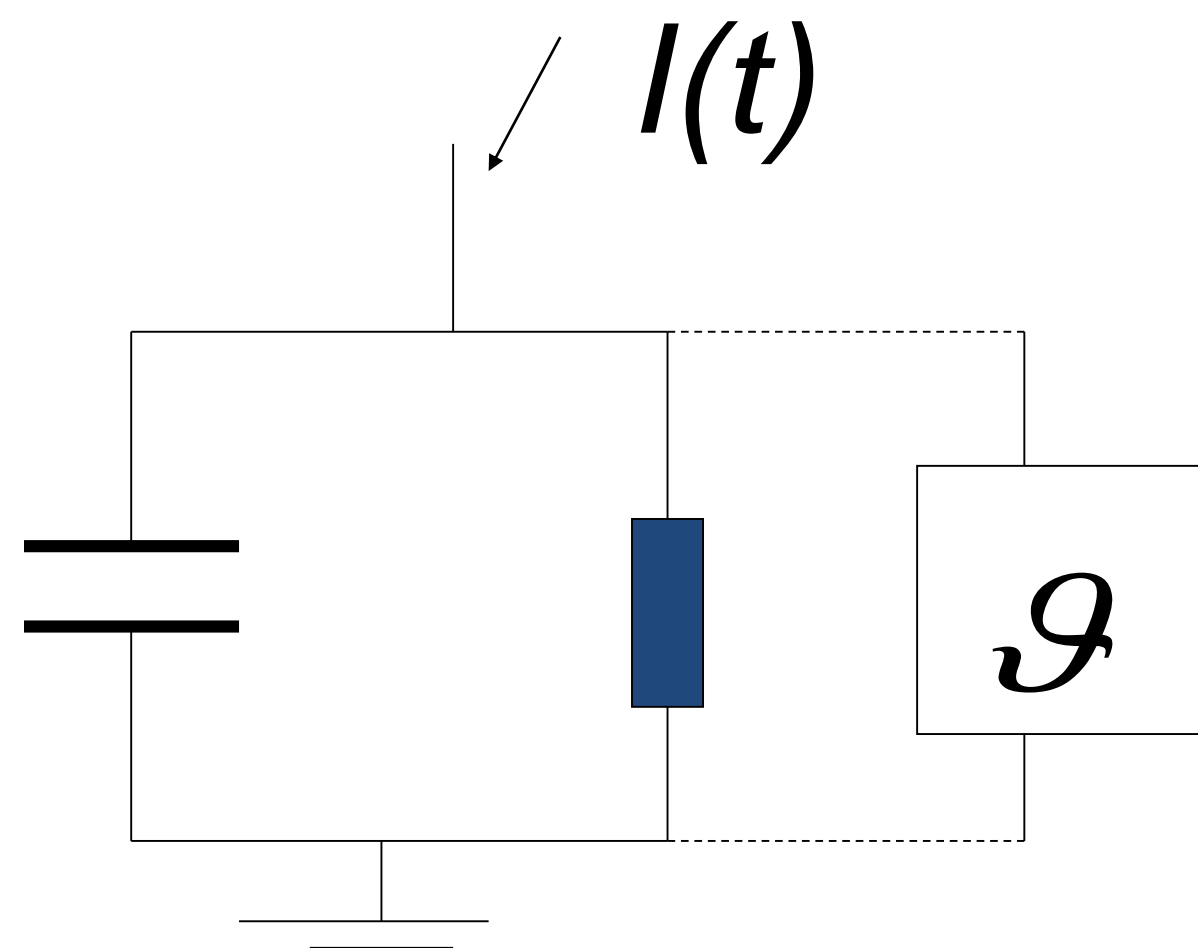
- (i) $x(t) = x_c \exp(t / \tau)$
- (ii) $x(t) = x_c \exp(-t / \tau)$
- (iii) $x(t) = x_c [1 - \exp(-t / \tau)]$
- (iv) $x(t) = 0.5x_c [1 + \exp(-t / \tau)]$

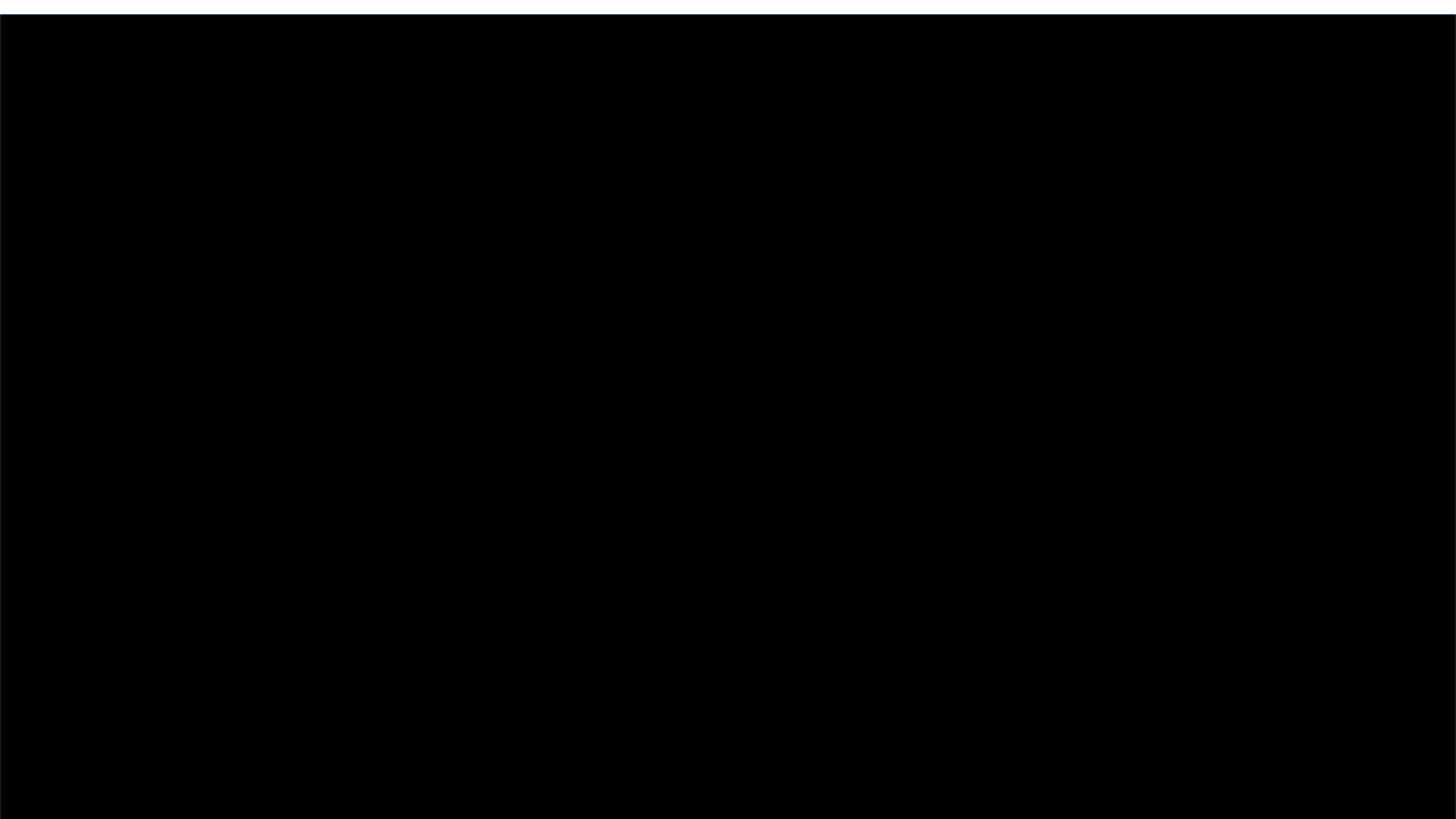
You will have to use the results: response to constant input/step input again and again

Neuronal Dynamics – 1.3 Leaky Integrate-and-Fire Model



CONSTANT input/step input



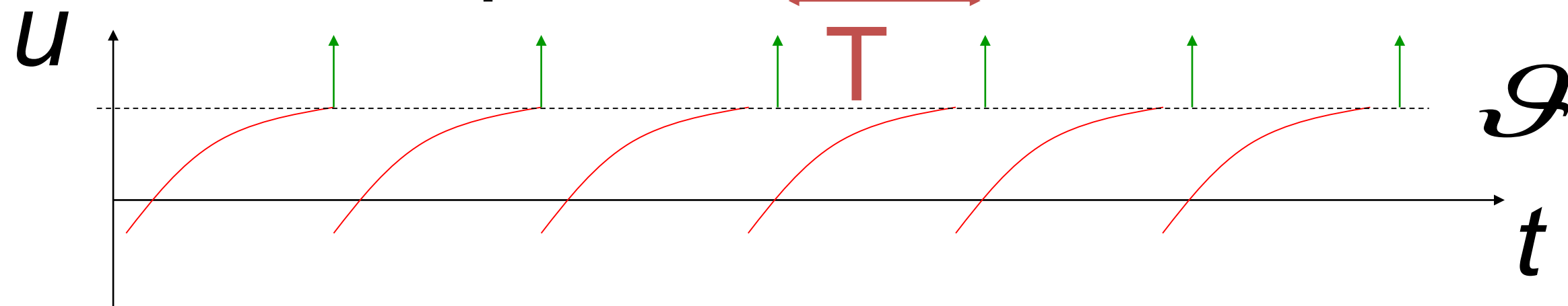


Leaky Integrate-and-Fire Model (LIF)

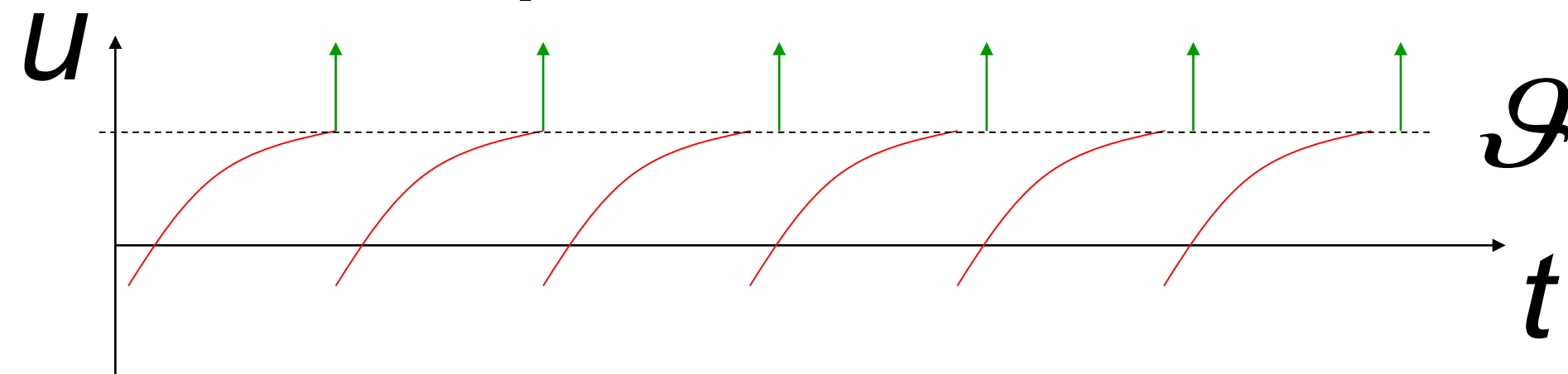
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI_0$$

LIF
If $u(t) = \mathcal{G} \Rightarrow u \rightarrow u_r$

Repetitive, current I_0



Repetitive, current $I_1 > I_0$

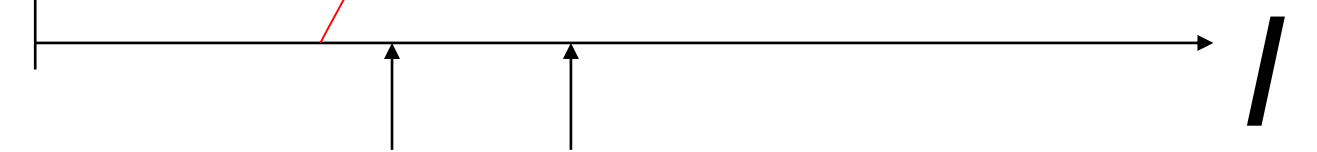


'Firing'

$1/T$

frequency-current
relation

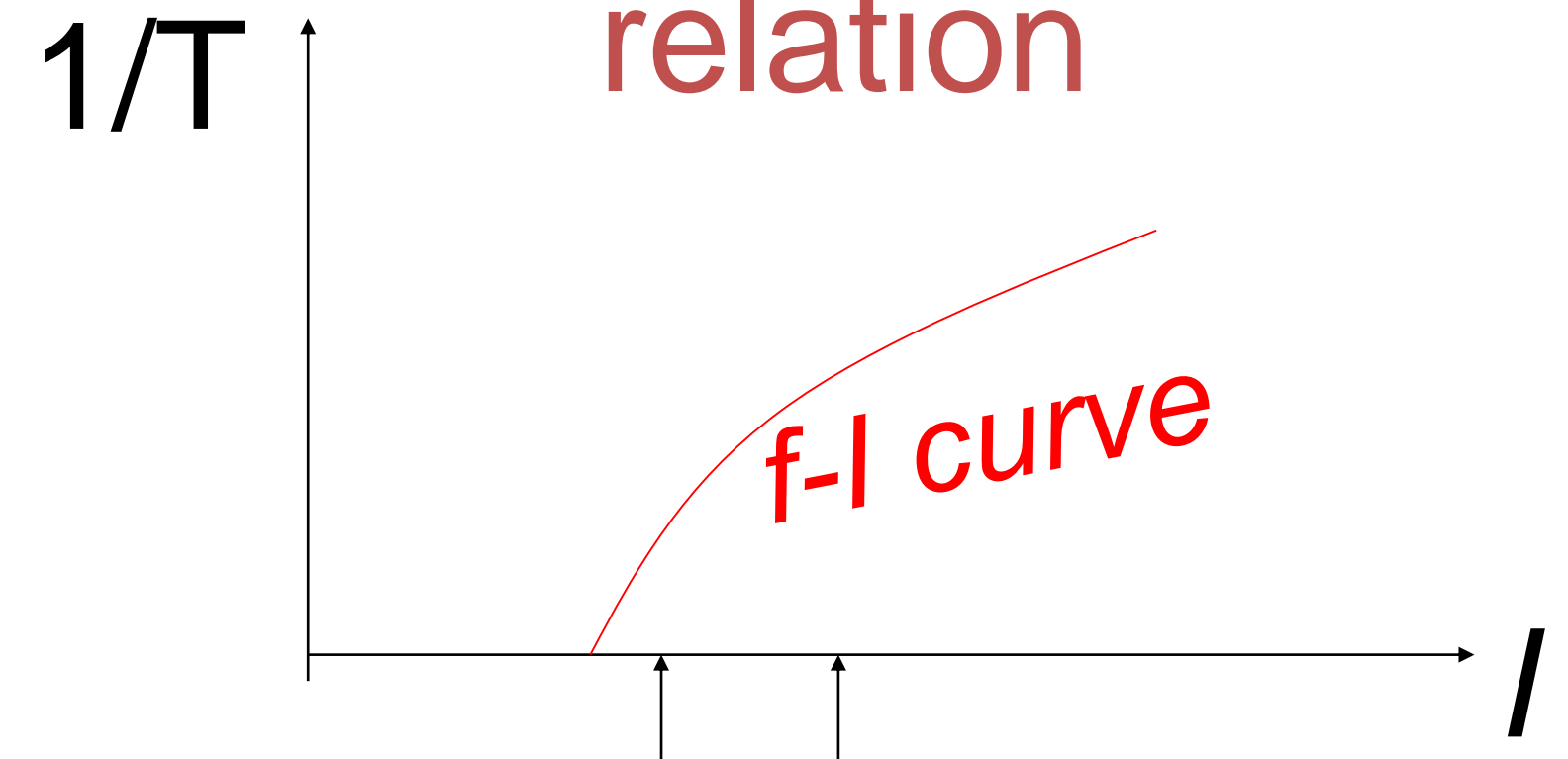
f-I curve



Neuronal Dynamics – First week, Exercise 2

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

frequency-current
relation



EXERCISE 2 NOW: Leaky Integrate-and-fire Model (LIF)

LIF $\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI_0$ If firing: $u \rightarrow u_r$

Exercise!

Calculate the

interspike interval T

for constant input I .

Firing rate is $f=1/T$.

Write f as a function of I .

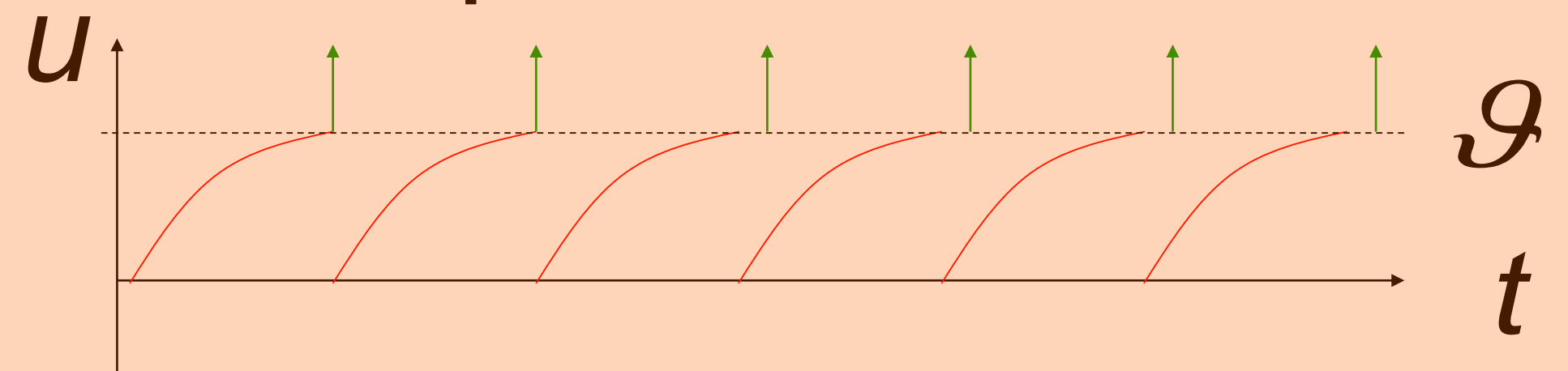
What is the

frequency-current curve

$f=g(I)$ of the LIF?

assume : $u_r = u_{rest}$

repetitive



**Start Exerc. at 10:53.
Next lecture at
11:15**

Exercise 2: Integrate-and-fire model

Consider the model of Eq. (1) with a threshold at $u = \vartheta > u_{\text{rest}}$. If the membrane potential reaches the threshold, the neuron is said to fire and the membrane potential is reset to u_{rest} . The injected current is a step of magnitude I_0 :

$$I(t) = \begin{cases} 0 & t \leq t_0 \\ I_0 & t > t_0 \end{cases}$$

- 2.1 What is the minimal current to reach the threshold, assuming $u(t = 0) = u_{\text{rest}}$?
- 2.2 At what time will the voltage first reach the threshold?
- 2.3 Calculate the firing frequency f as a function of I_0 .

The function $g(I_0)$ which gives the firing frequency as a function of the constant applied current is called gain function.

Summary of Section 1.3.

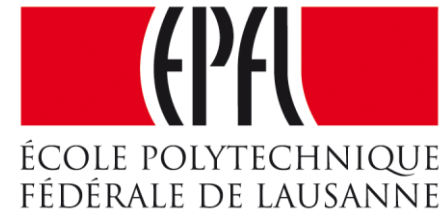
The leaky integrate-and-fire neuron model is the combination of a passive membrane (linear differential equation) with a threshold.

The moment when the potential hits the threshold defines the firing time of a spike. Immediately after firing the voltage is reset to a lower value (not necessarily to the resting potential).

For the leaky integrate-and-fire model, the gain function (frequency of firing for constant input, as a function of input strength) can be calculated analytically. The firing frequency decreases with decreasing input. If the constant input is below a critical value no firing can occur. This value defines the rheobase current threshold.

For the leaky integrate-and-fire model the rheobase current threshold can be predicted from the voltage threshold and the model parameters R and C .

Week 1 – part 4: Generalized Integrate-and-Fire Model



Biological Modeling of Neural Networks

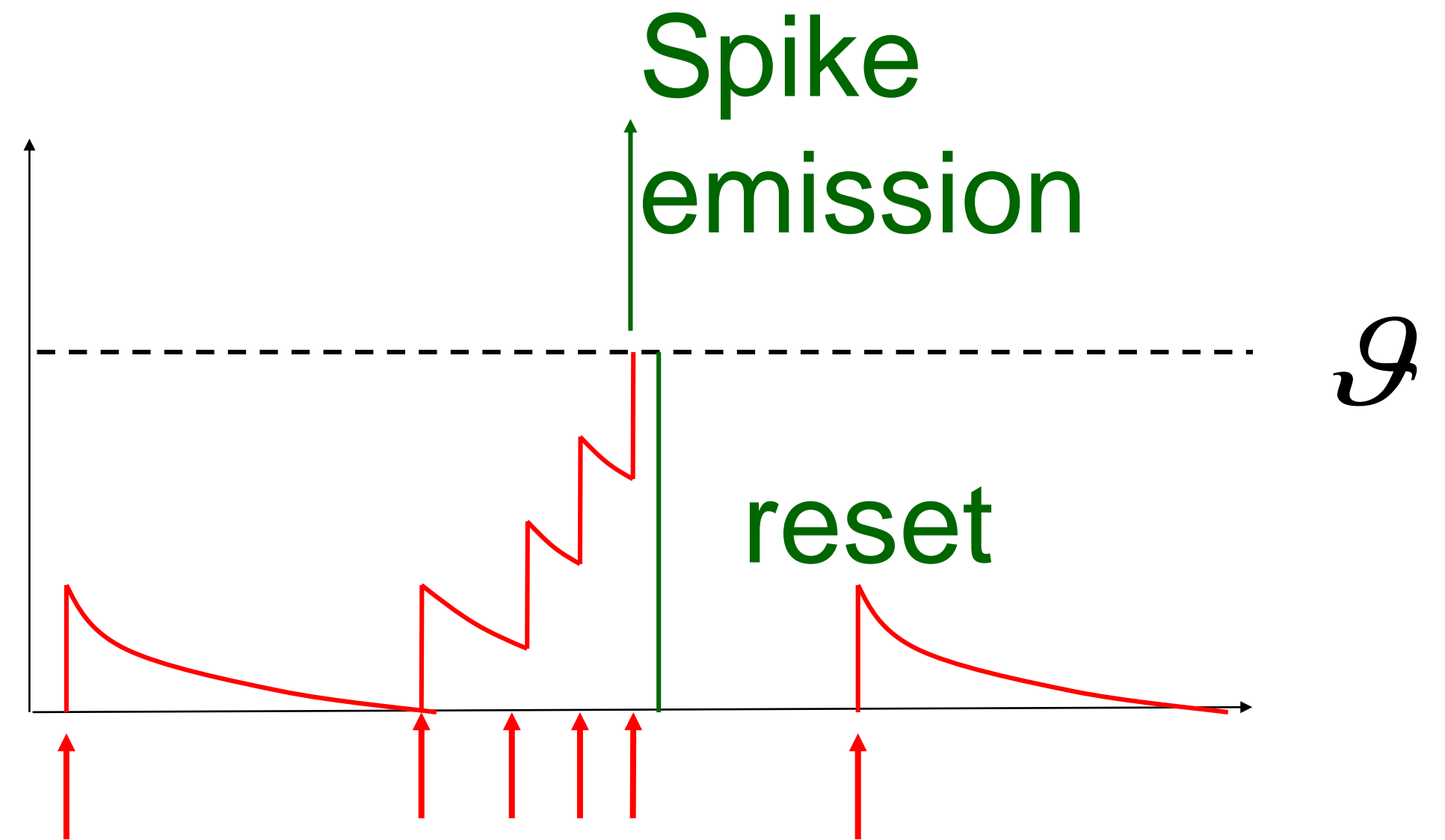
Week 1 – neurons and mathematics:
a first simple neuron model

Wulfram Gerstner

EPFL, Lausanne, Switzerland

- ✓ 1.1 Neurons and Synapses:
Overview
- ✓ 1.2 The Passive Membrane
 - Linear circuit
 - Dirac delta-function
- ✓ 1.3 Leaky Integrate-and-Fire Model
- 1.4 Generalized Integrate-and-Fire Model
- 1.5. Quality of Integrate-and-Fire Models

Neuronal Dynamics – 1.4. Generalized Integrate-and-Fire



Integrate-and-fire model

LIF: linear + threshold

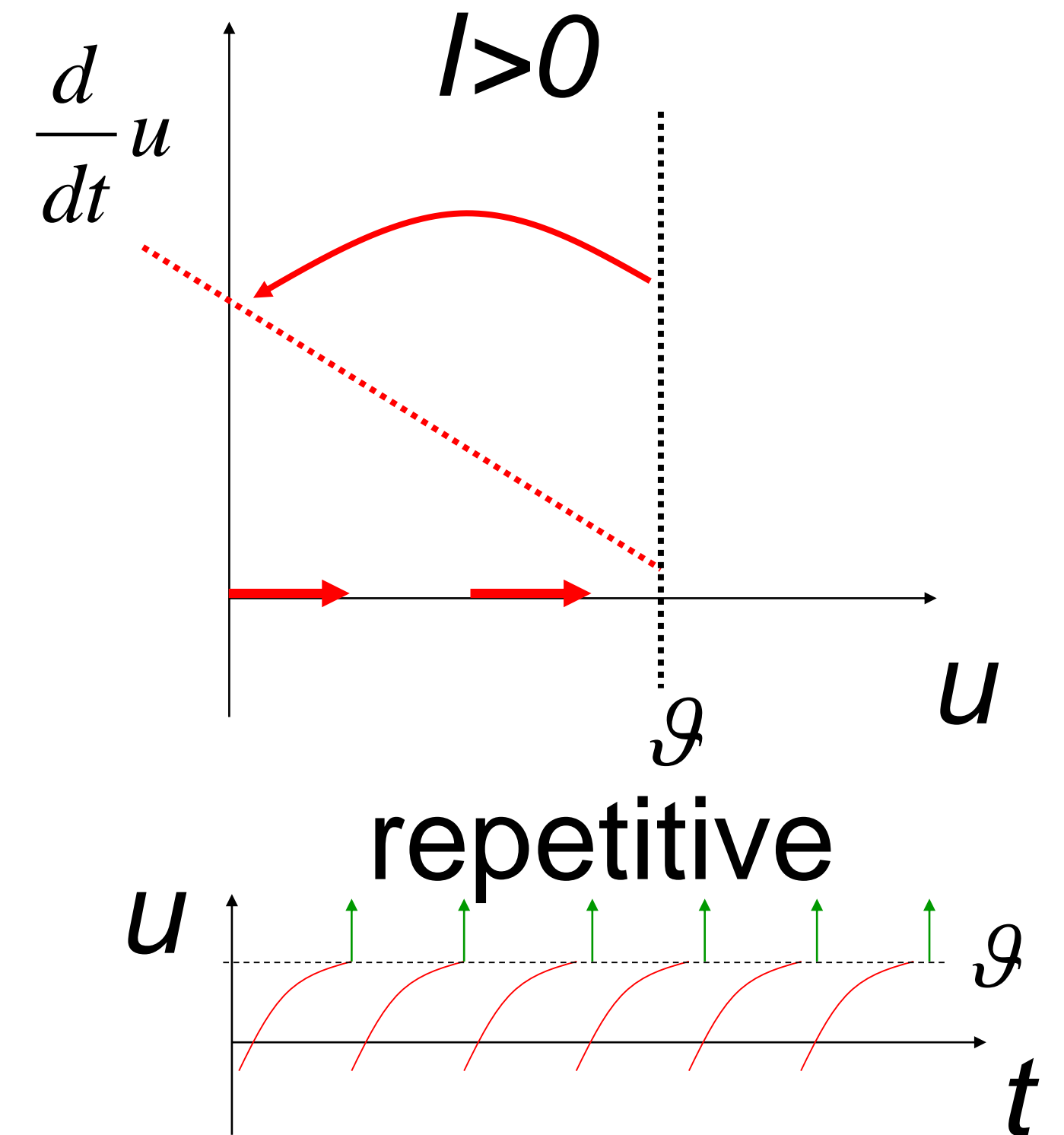
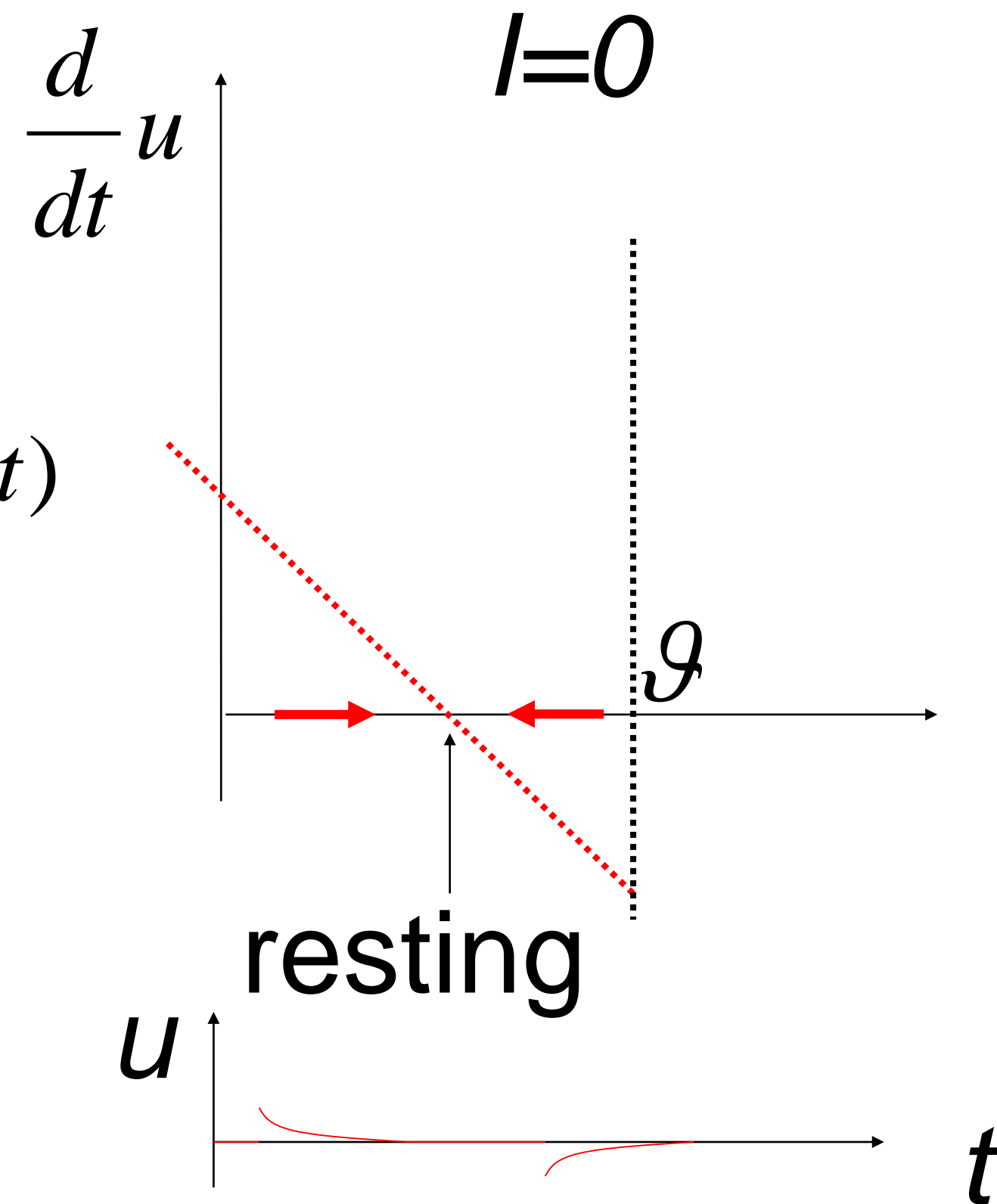
Neuronal Dynamics – 1.4. Leaky Integrate-and Fire revisited

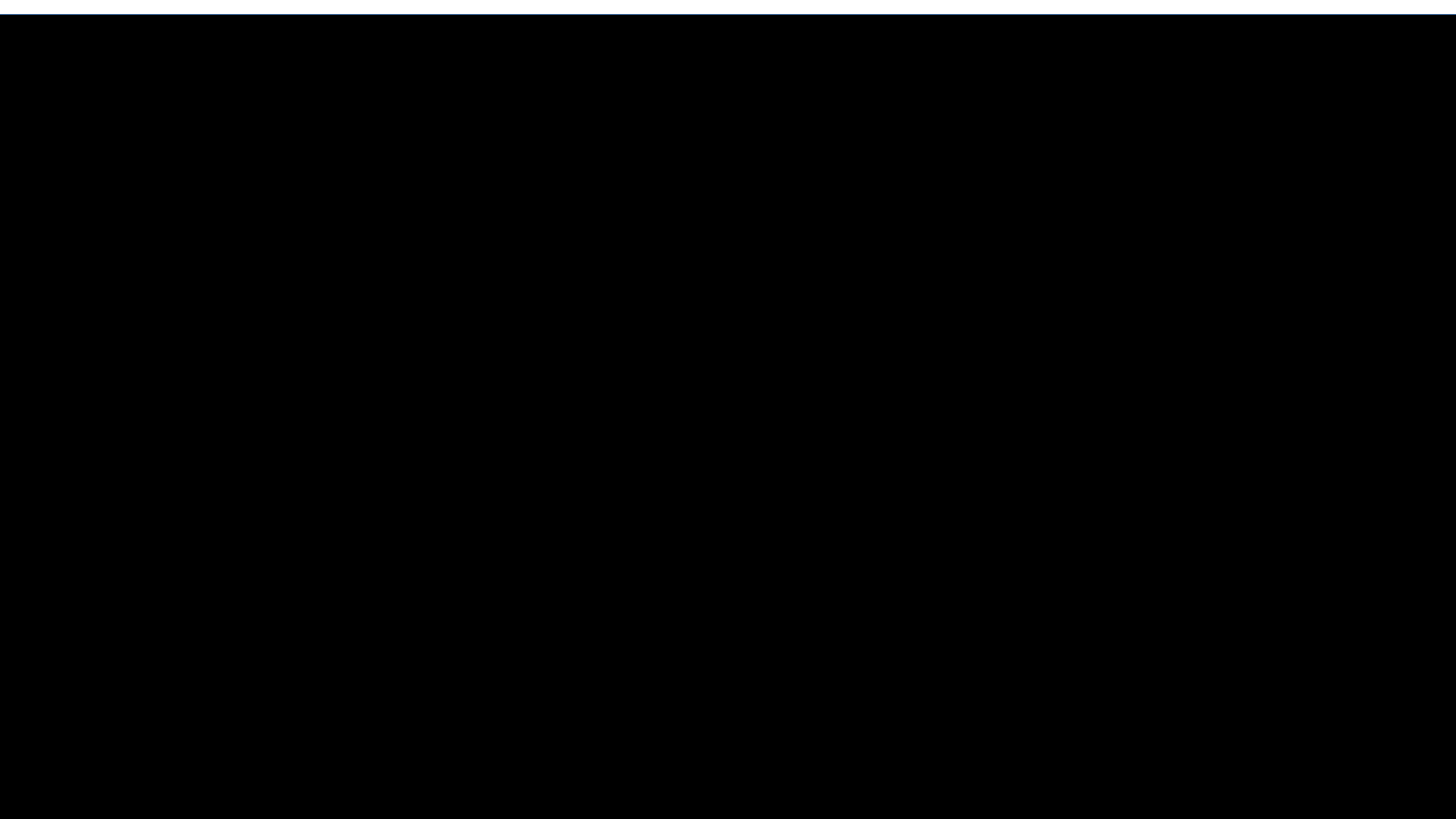
LIF

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

If firing:

$$u \rightarrow u_r$$





Neuronal Dynamics – 1.4. Nonlinear Integrate-and-Fire

LIF

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

NLIF

$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t)$$

If firing:

$$u \rightarrow u_{reset}$$

Neuronal Dynamics – 1.4. Nonlinear Integrate-and-Fire

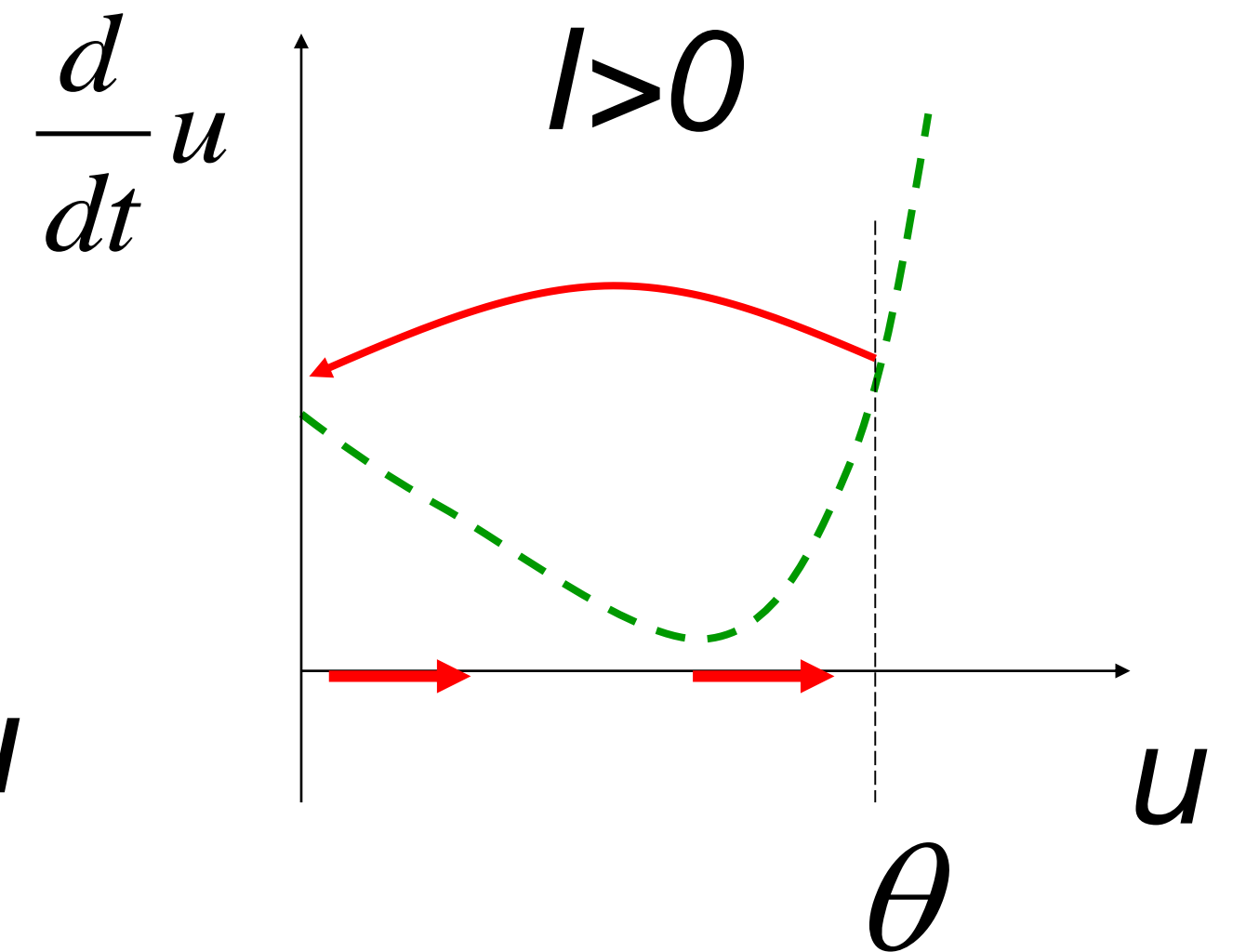
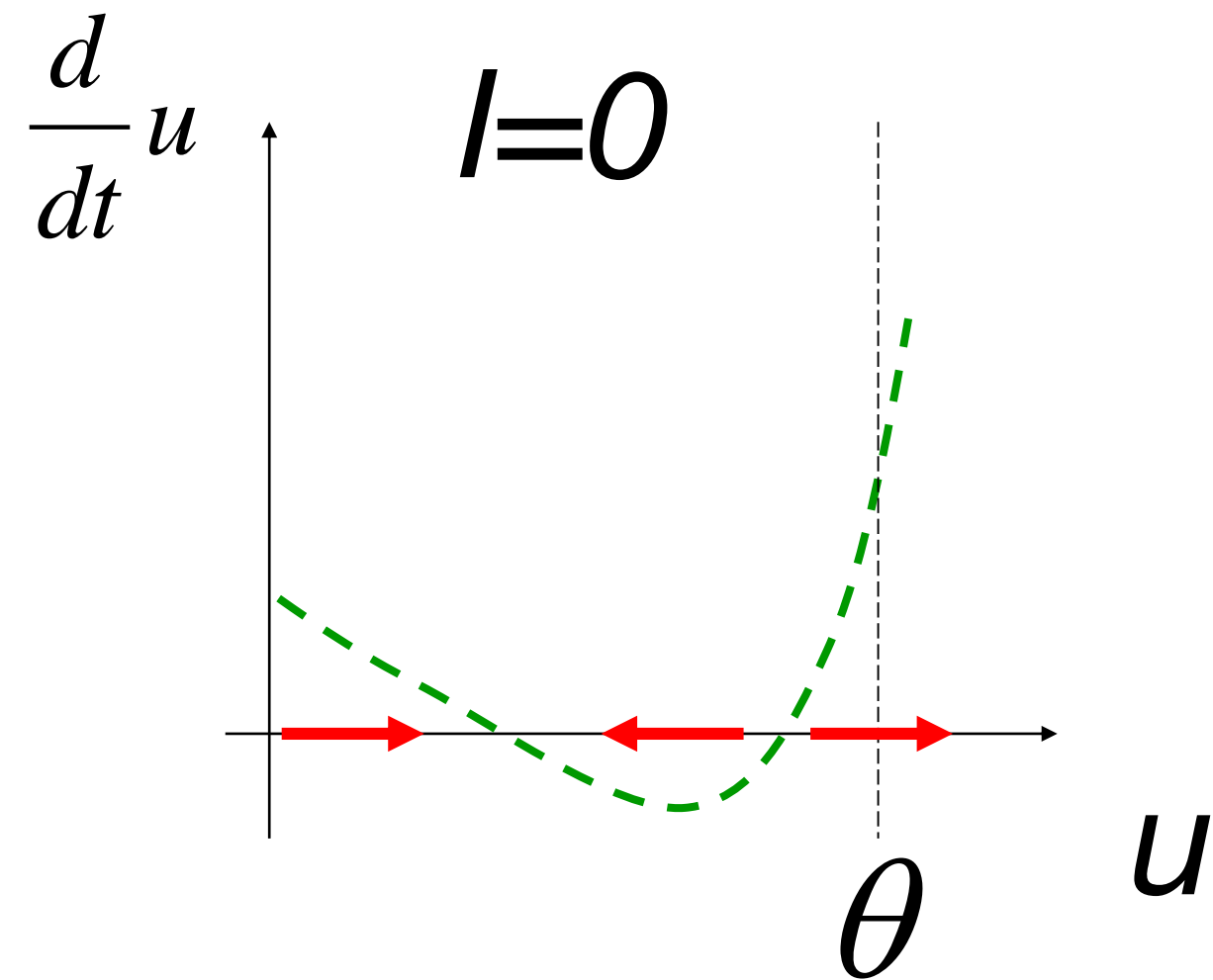
Nonlinear Integrate-and-Fire

NLIF

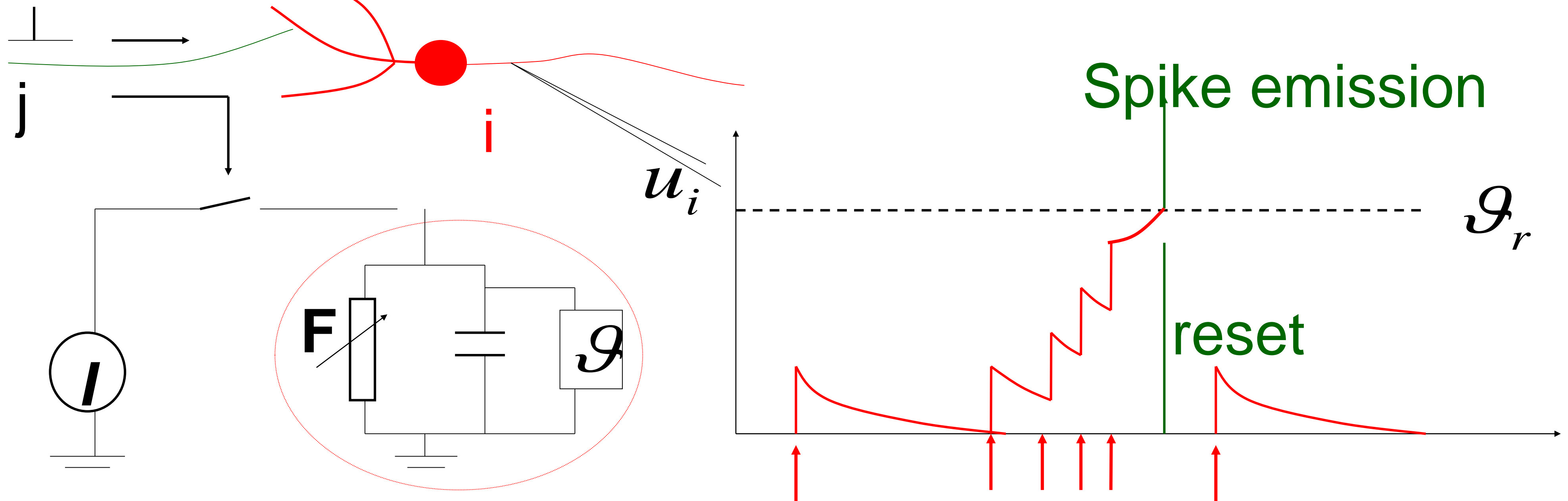
$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t)$$

firing: $u(t) = \theta \Rightarrow$

$$u \rightarrow u_r$$



Nonlinear Integrate-and-fire Model

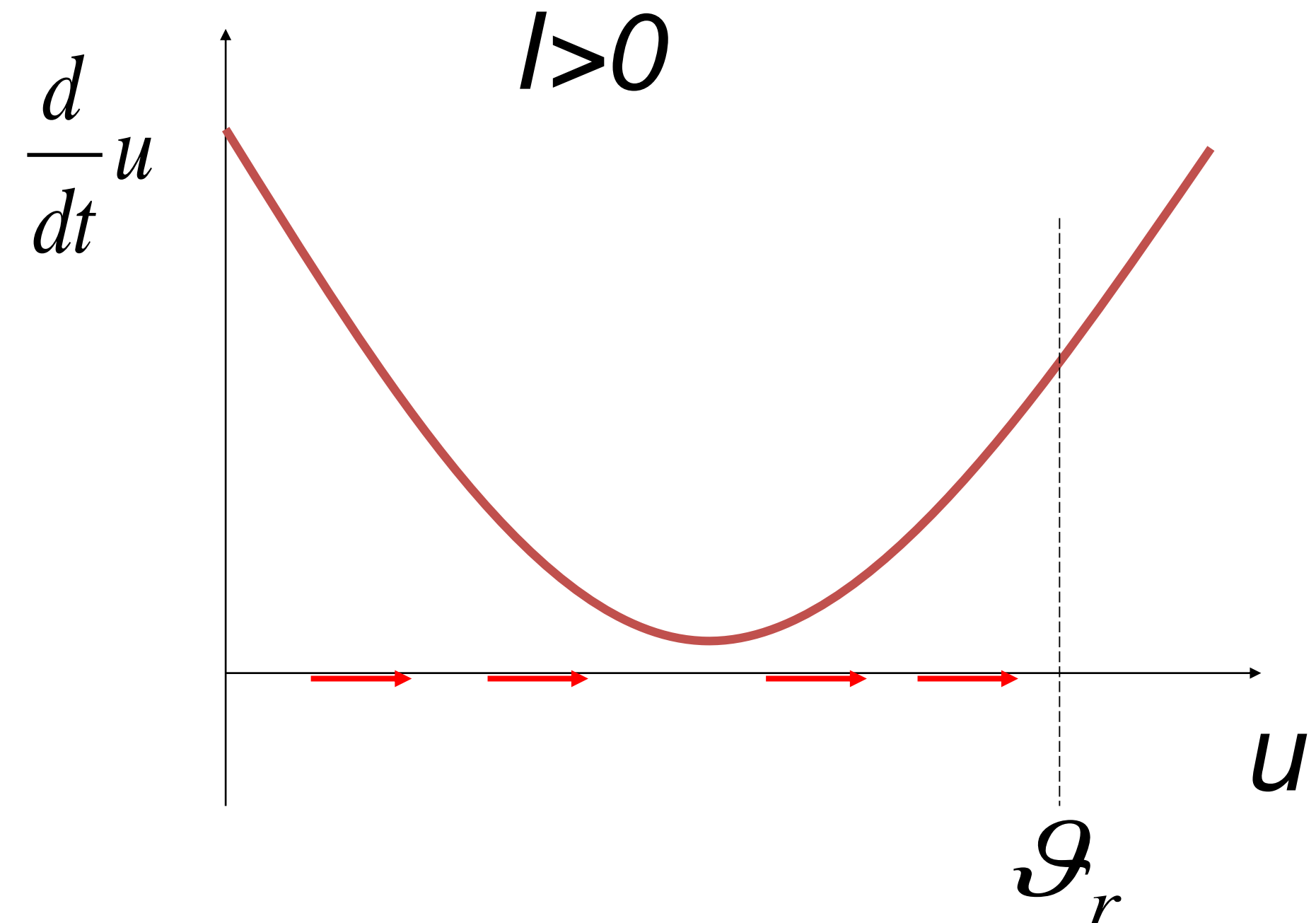
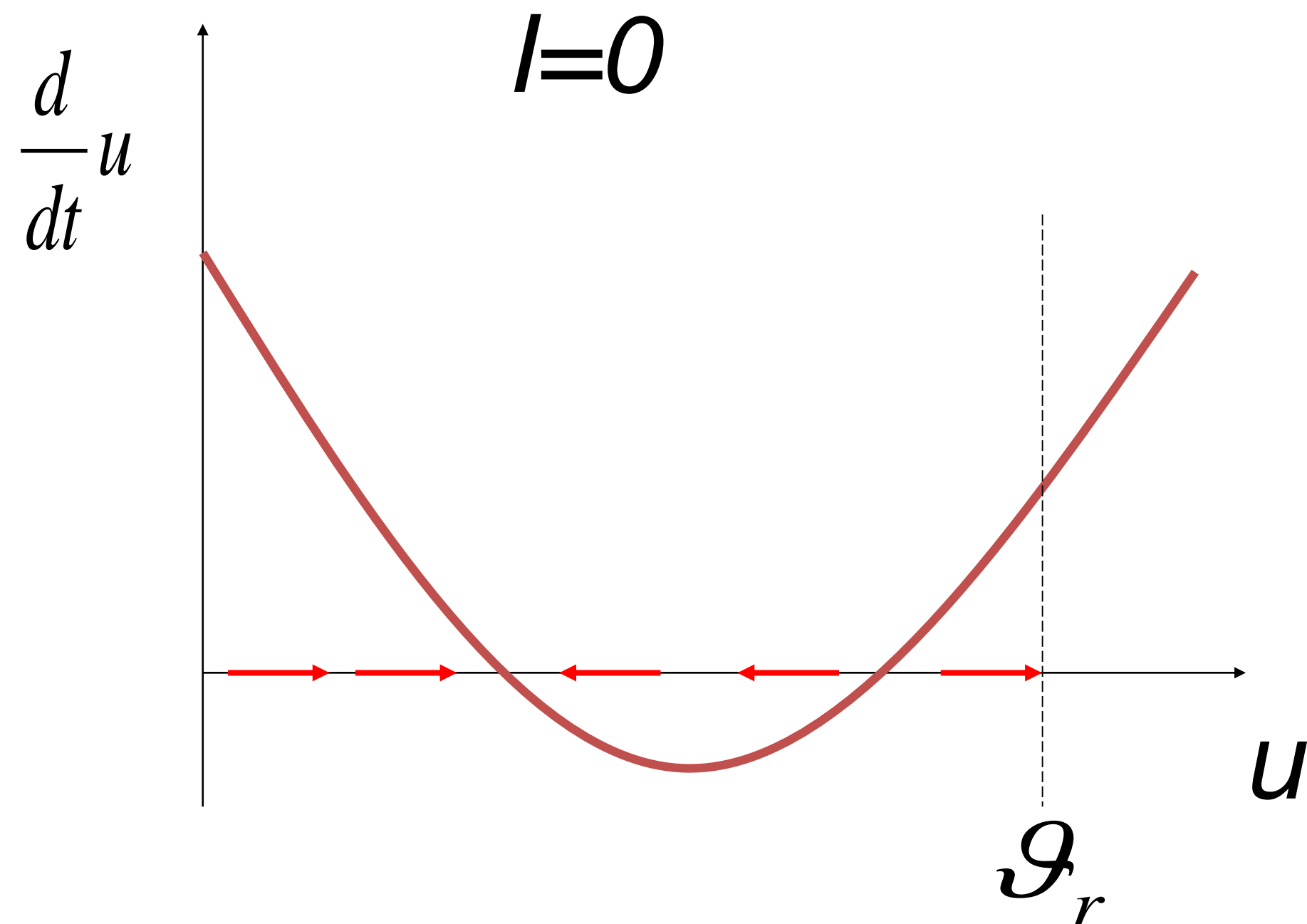


$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t)$$

NONlinear

$$u(t) = \mathcal{G}_r \Rightarrow \text{Fire+reset threshold}$$

Nonlinear Integrate-and-fire Model

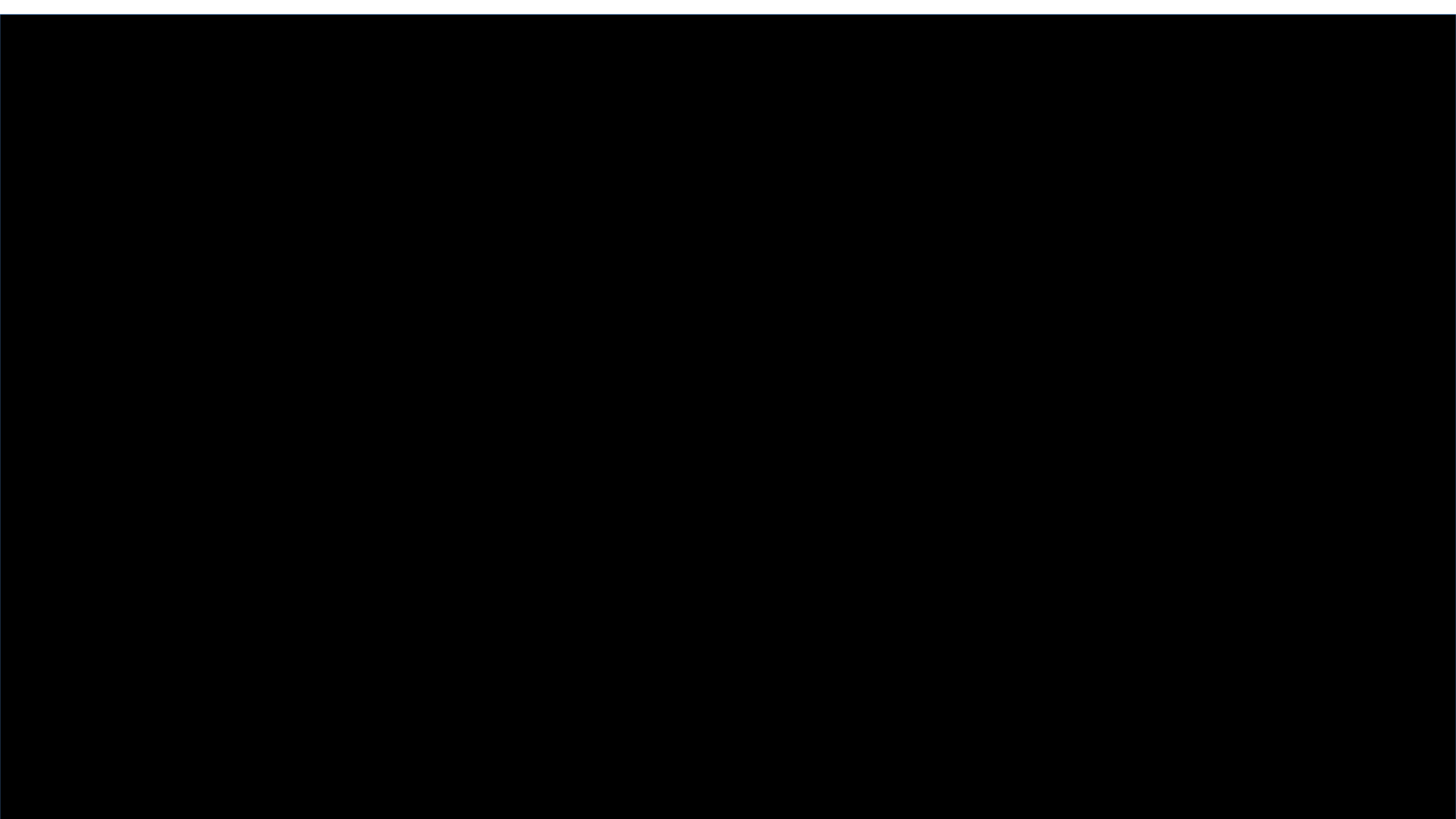


$$\tau \cdot \frac{d}{dt}u = F(u) + RI(t) \quad \text{NONlinear}$$

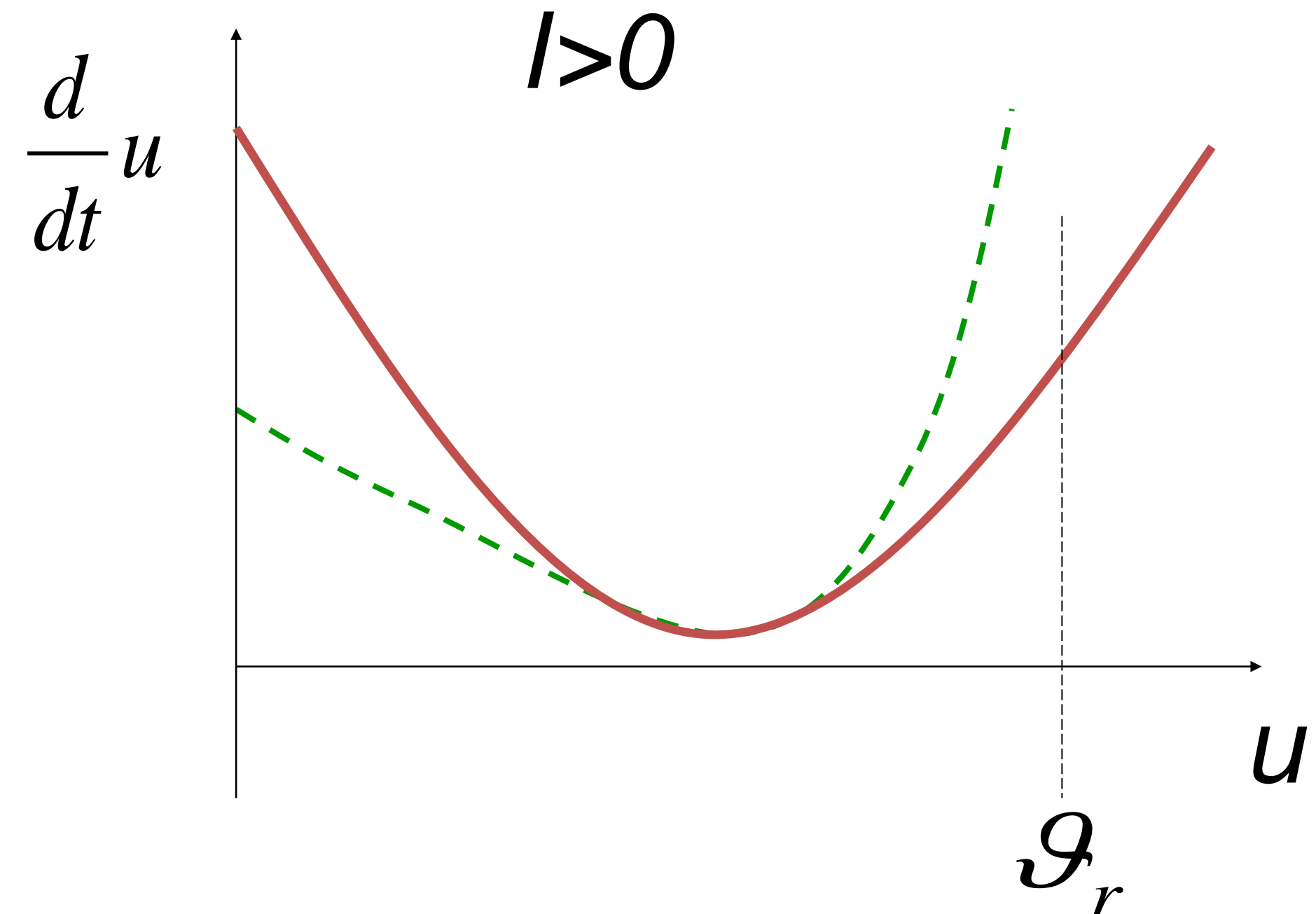
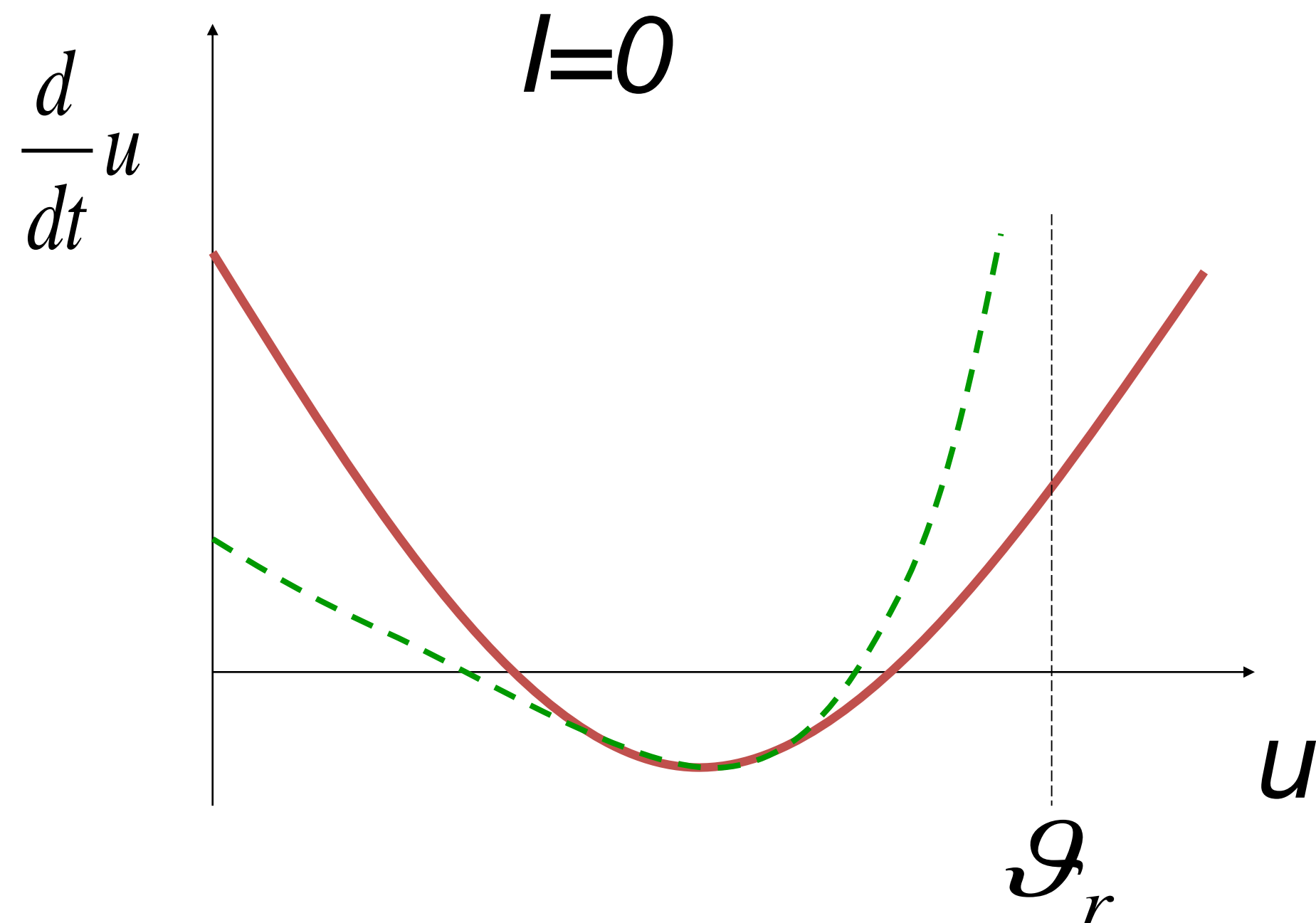
$$u(t) = \mathcal{G}_r \Rightarrow \text{Fire+reset threshold}$$

Quadratic I&F:

$$F(u) = c_2(u - c_1)^2 + c_0$$



Nonlinear Integrate-and-fire Model



$$\tau \cdot \frac{d}{dt}u = F(u) + RI(t)$$

$$u(t) = \mathcal{G}_r \Rightarrow \text{Fire+reset}$$

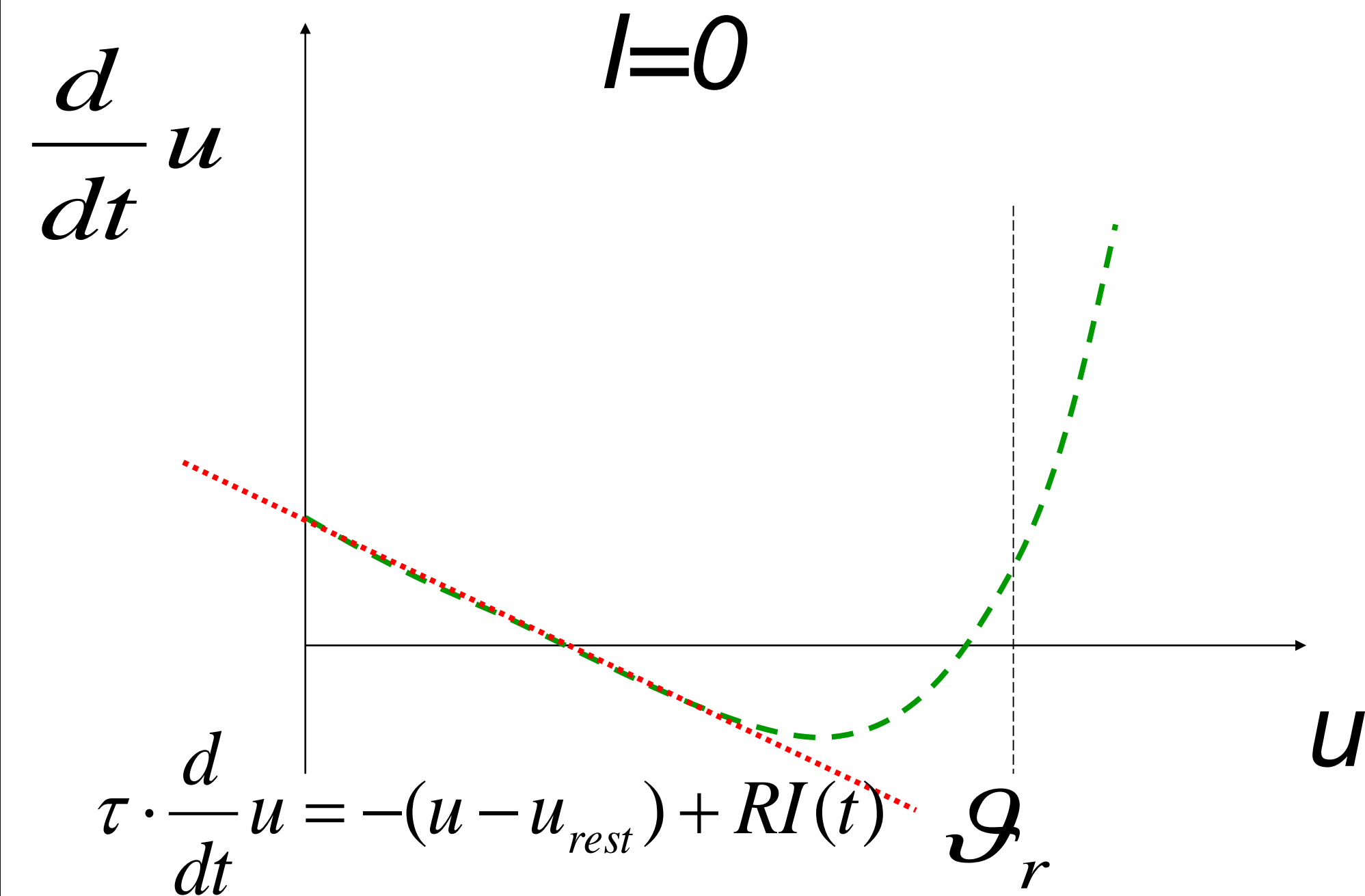
Quadratic I&F:

$$F(u) = c_2(u - c_1)^2 + c_0$$

exponential I&F:

$$F(u) = -(u - u_{rest}) + c_0 \exp(u - \mathcal{G})$$

Nonlinear Integrate-and-fire Model



$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t) \quad \text{NONlinear}$$

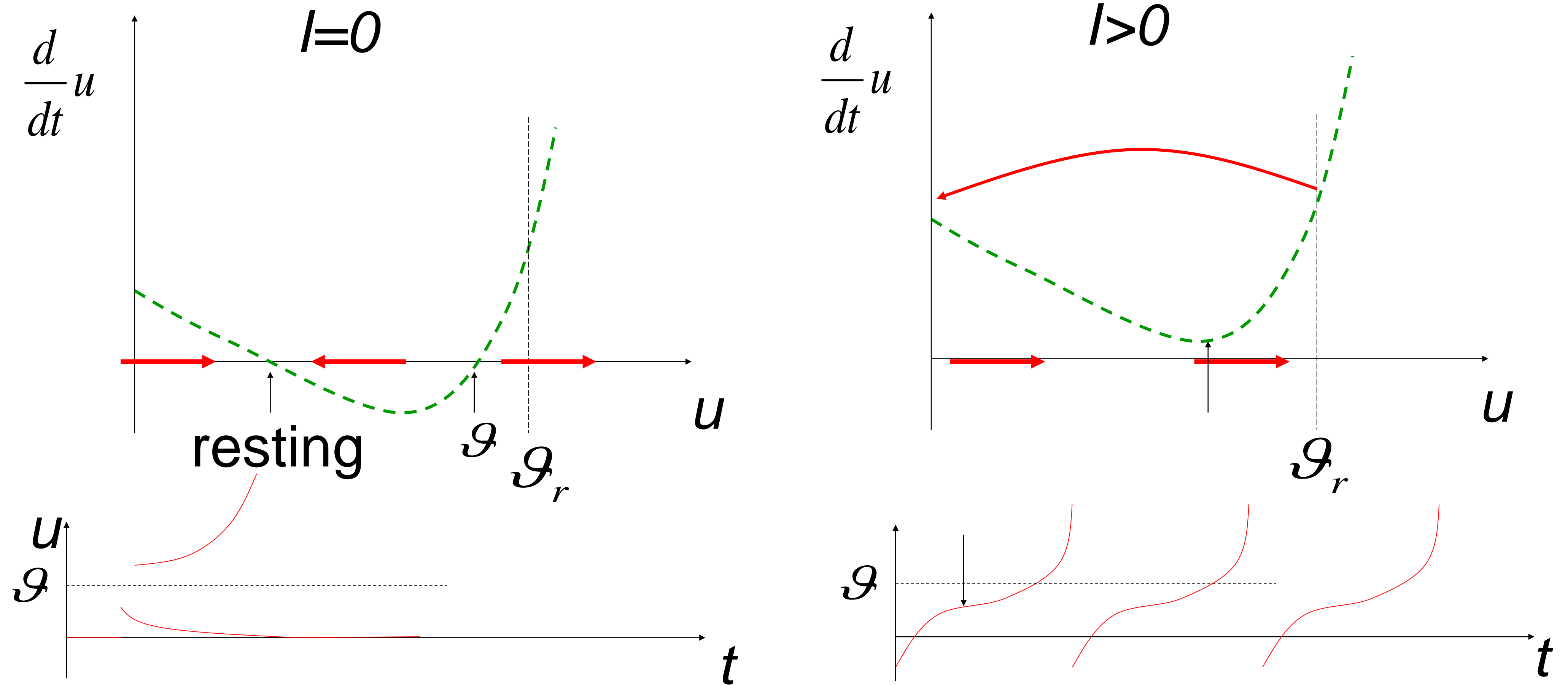
$$u(t) = \mathcal{G}_r \implies \text{Fire+reset threshold}$$

exponential I&F:

$$F(u) = -(u - u_{rest}) + c_0 \exp(u - \mathcal{G})$$

Nonlinear Integrate-and-fire Model

Where is the firing threshold?



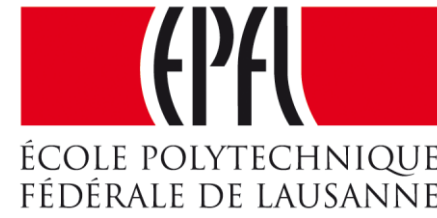
$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t)$$

Summary of Section 1.4.

Nonlinear integrate-and-fire models with a nonlinear function $f(u)$ of the voltage u are excellent models of spiking behavior. The exponential integrate-and-fire (EIF) model is linear for low voltage, but combines it with an exponential nonlinearity for large voltages. The quadratic integrate-and-fire (QIF) is symmetric for low and large voltages. The LIF has a linear function $f(u) = u - u_{rest}$. All generalized integrate-and-fire models have a voltage reset after a spike. For constant input, the zero-crossings of the function $f(u)$ define stationary states.

We distinguish two important stimulation paradigms: (i) stimulation with short current pulses (Dirac delta pulses) – such pulses correspond to an initial condition on the voltage axis; (ii) stimulation with constant current – such stimuli correspond to a vertical shift of the function $f(u)$. We find in both EIF and QIF that the minimal current for spike initiation depends on the stimulation protocol.

Week 1 – part 5: How good are Integrate-and-Fire Model?



Biological Modeling of Neural Networks

Week 1 – neurons and mathematics:
a first simple neuron model

Wulfram Gerstner

EPFL, Lausanne, Switzerland

↓ 1.1 Neurons and Synapses:

Overview

1.2 The Passive Membrane

- Linear circuit
- Dirac delta-function

↓ 1.3 Leaky Integrate-and-Fire Model

↓ 1.4 Generalized Integrate-and-Fire Model

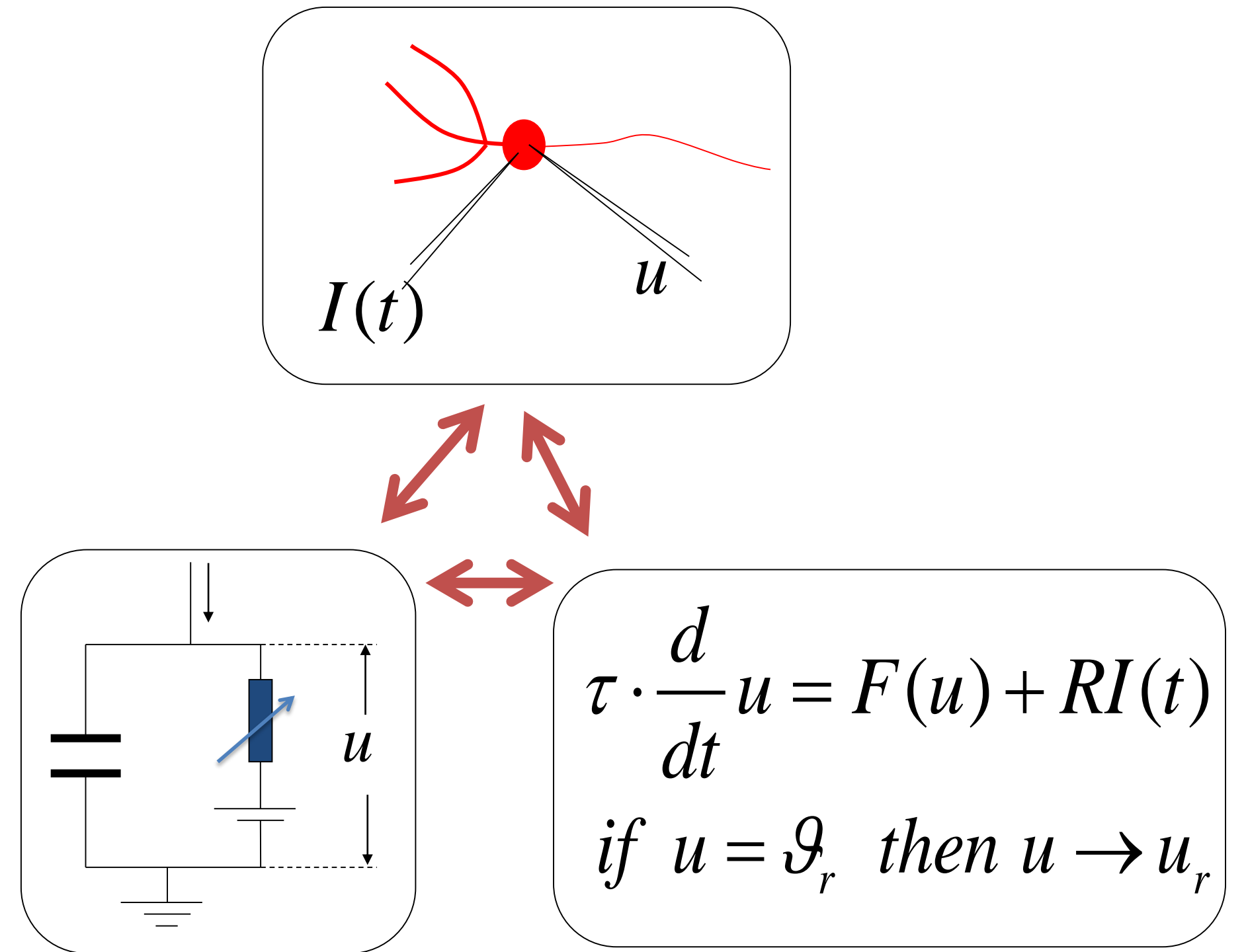
- where is the firing threshold?

1.5. Quality of Integrate-and-Fire Models

- Neuron models and experiments

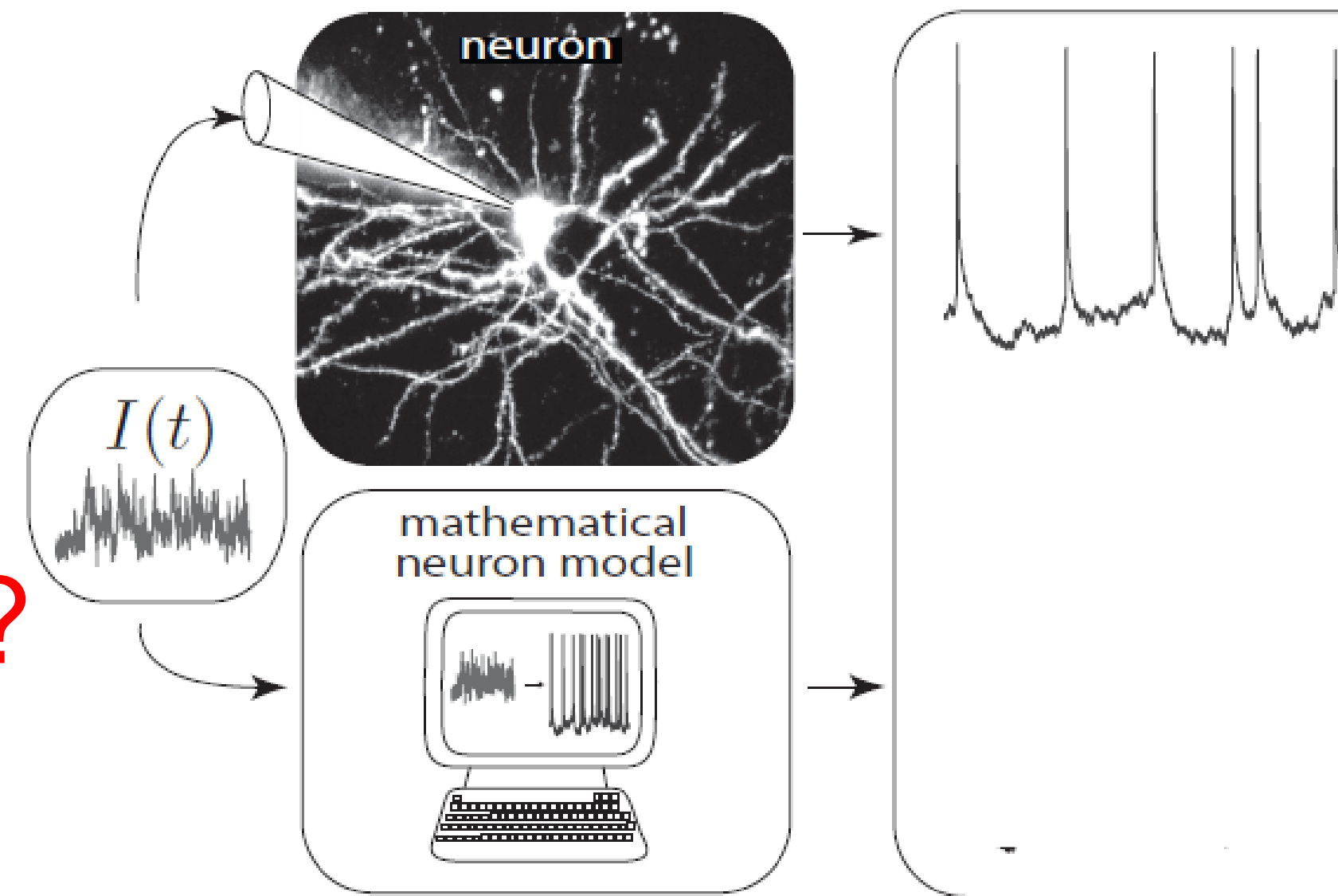
Neuronal Dynamics – 1.5. How good are integrate-and-fire models?

Can we compare neuron models
with experimental data?



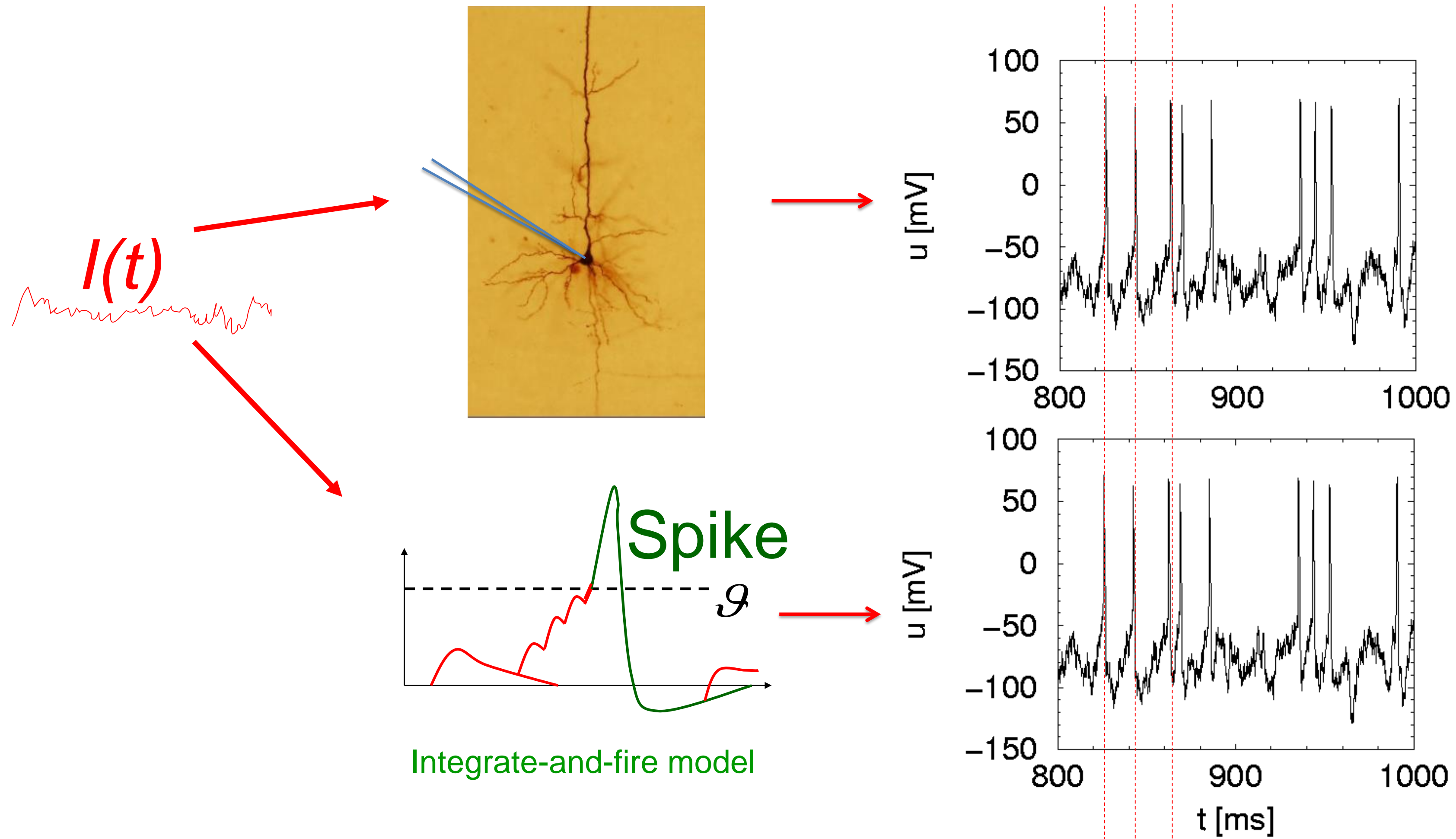
Neuronal Dynamics – 1.5. How good are integrate-and-fire models?

What is a good neuron model?

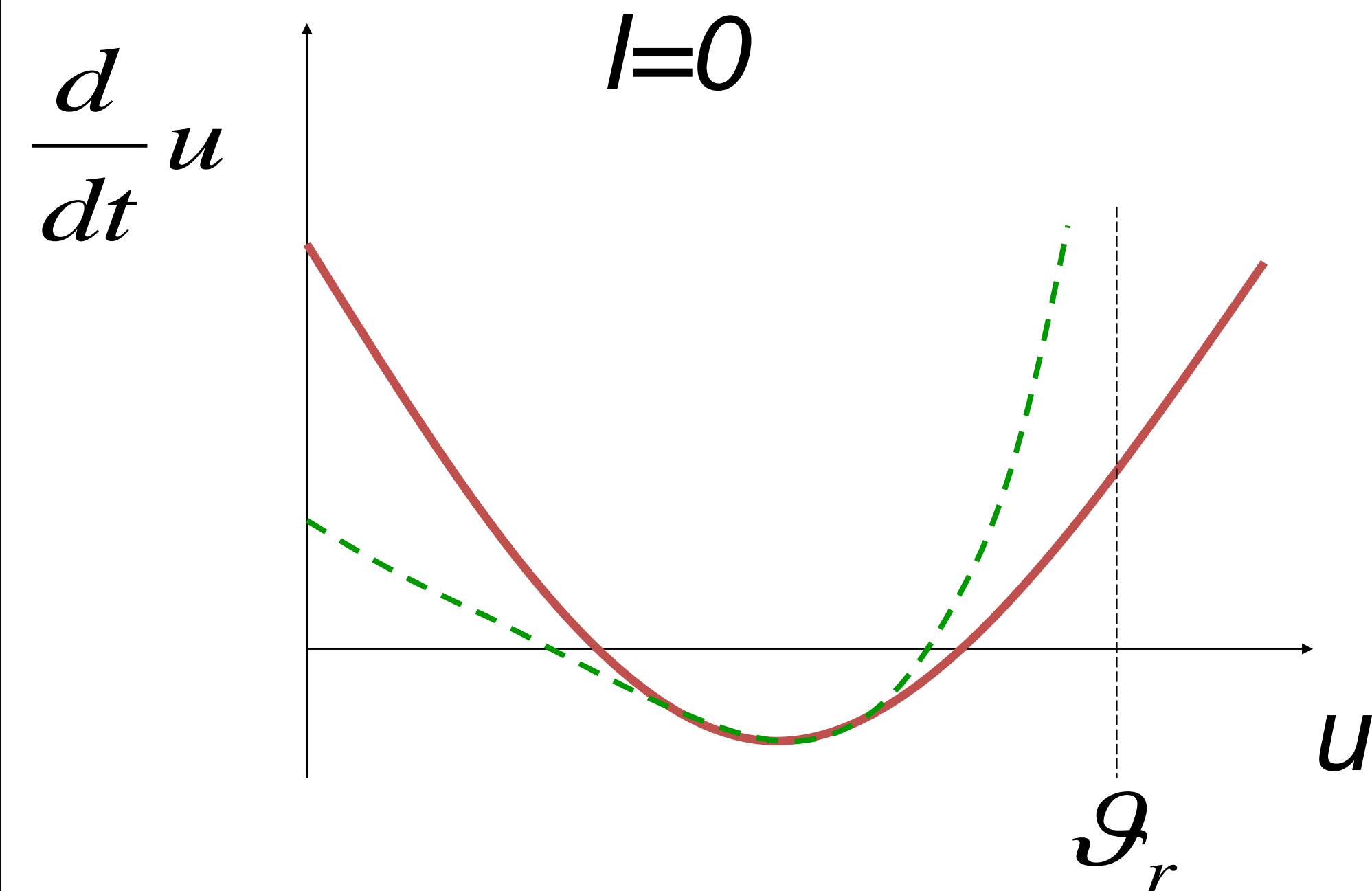


Can we compare neuron models
with experimental data?

Neuronal Dynamics – 1.5. How good are integrate-and-fire models?



Nonlinear Integrate-and-fire Model



Can we measure
the function $F(u)$?

$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t)$$

$$u(t) = \mathcal{G}_r \Rightarrow \text{Fire+reset}$$

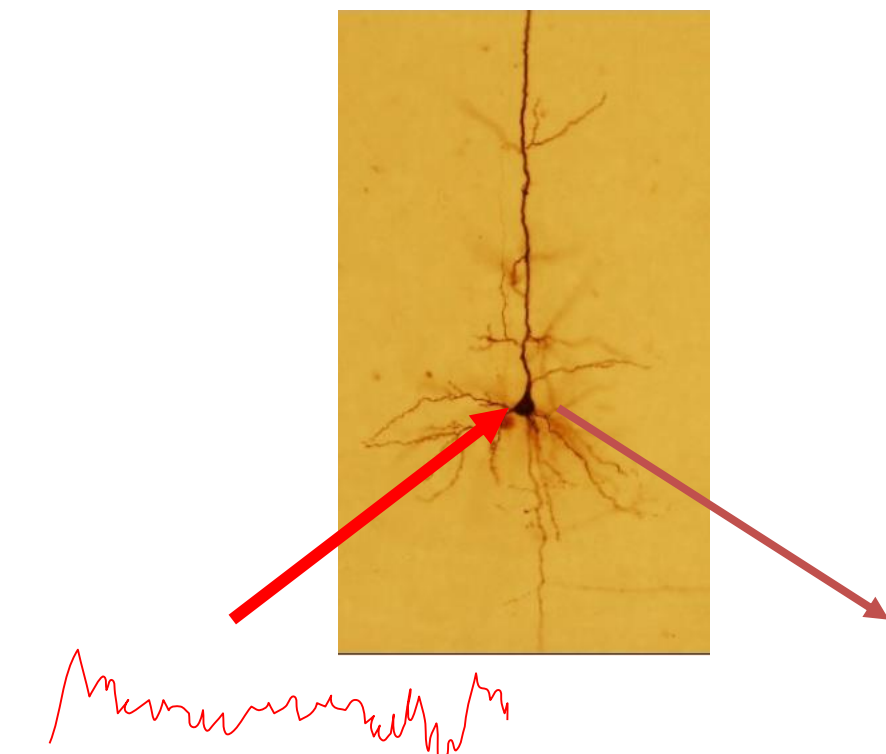
Quadratic I&F:

$$F(u) = c_2(u - c_1)^2 + c_0$$

exponential I&F:

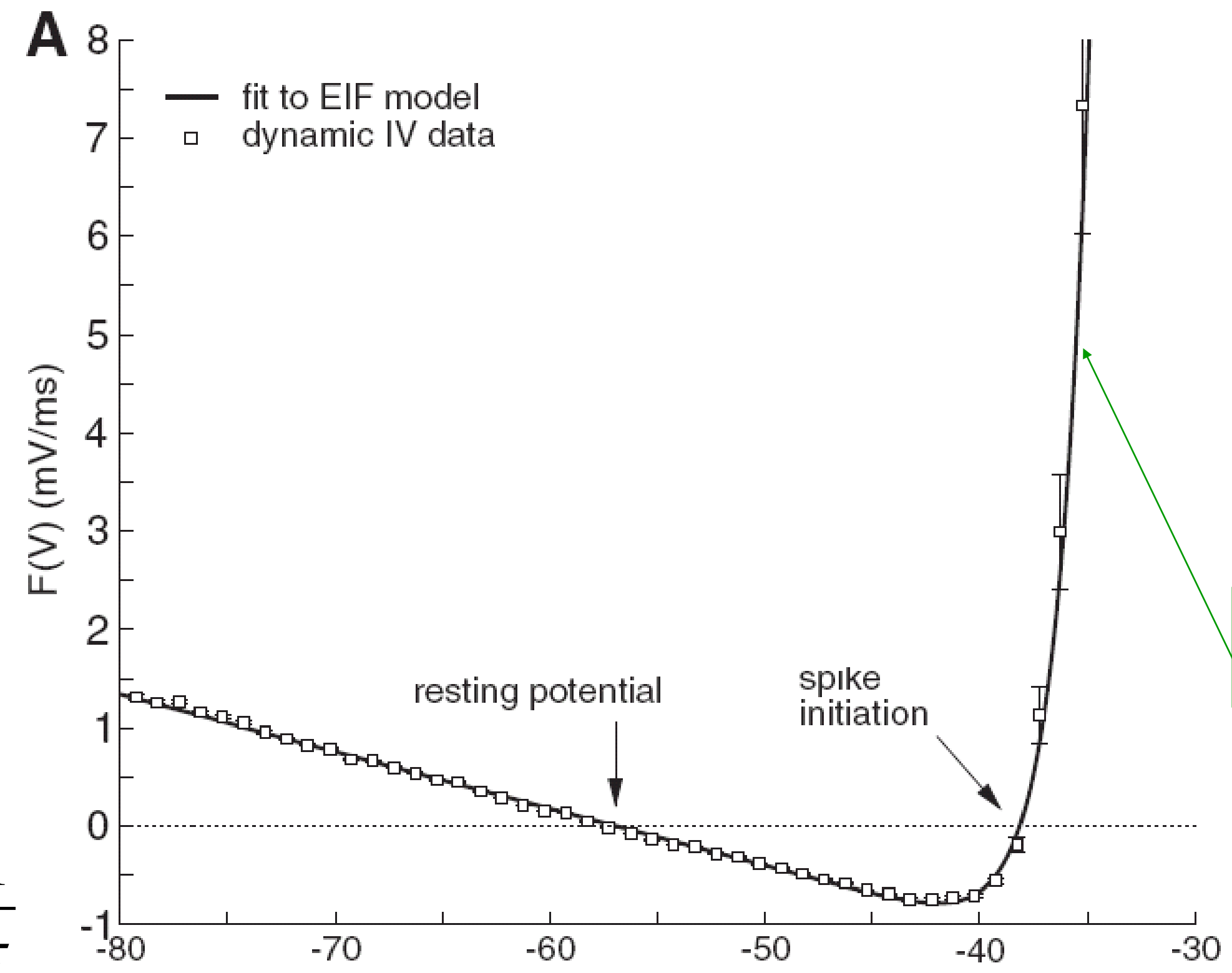
$$F(u) = -(u - u_{rest}) + c_0 \exp(u - \mathcal{G})$$

Neuronal Dynamics – 1.5. How good are integrate-and-fire models?



$I(t)$

$$\frac{du}{dt} - \frac{1}{C} I(t) = F(u) \frac{1}{\tau}$$

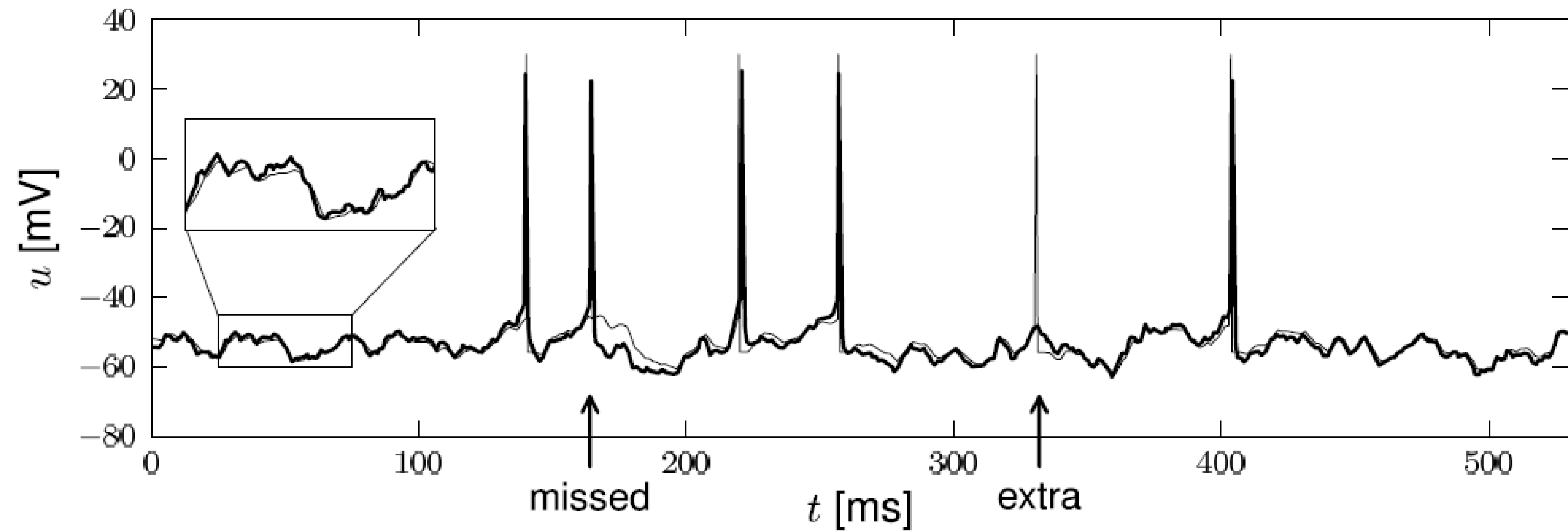
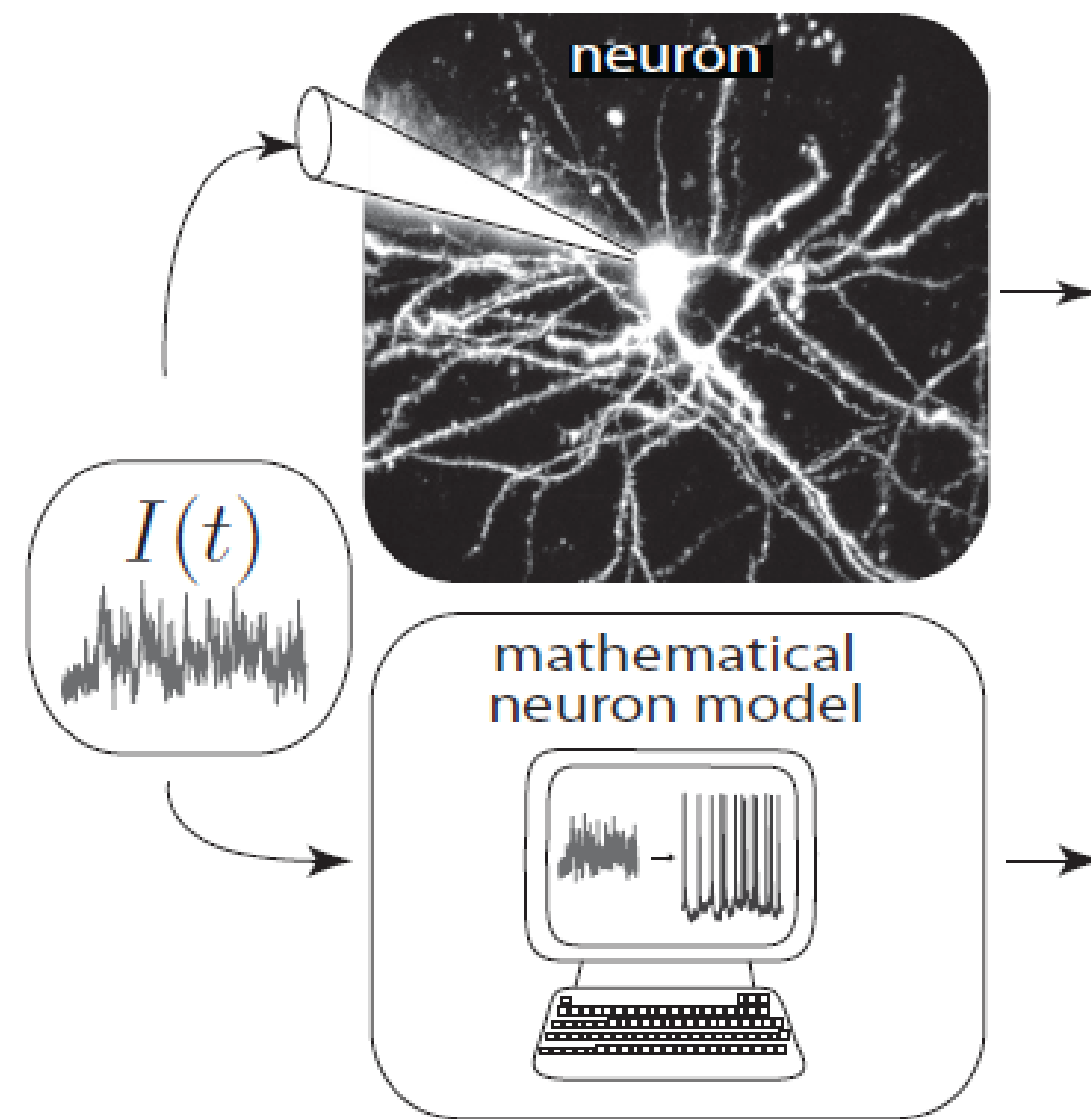


$$F(u) = -(u - u_{rest}) + \Delta \exp\left(\frac{u - \mathcal{G}}{\Delta}\right)$$

u [mV]

Badel et al., J. Neurophysiology 2008

Neuronal Dynamics – 1.5. How good are integrate-and-fire models?



Neuronal Dynamics – 1.5. How good are integrate-and-fire models?

Nonlinear integrate-and-fire models
are good

Mathematical description → prediction

Computer exercises:
Python

Need to add

- adaptation
- noise
- dendrites/synapses

Biological Modeling of Neural Networks

<http://neurondynamics.epfl.ch/>

Textbook:

Lecture today:

-Chapter 1

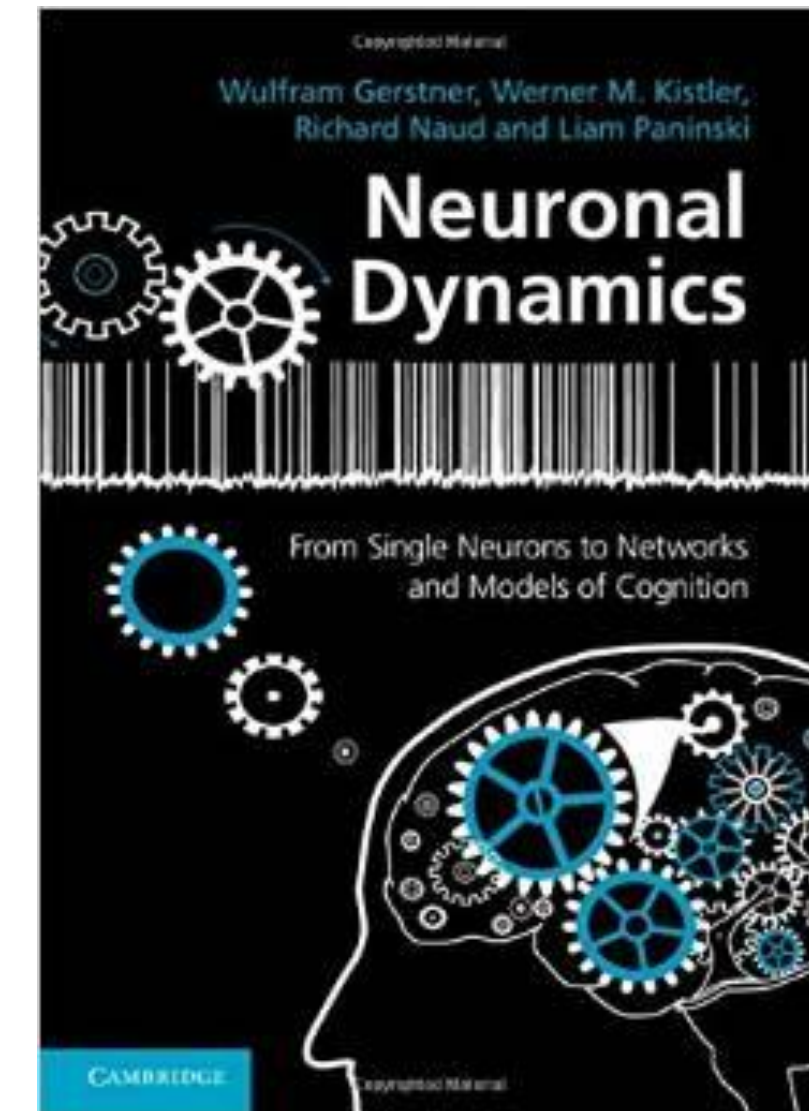
-Chapter 5

Exercises today:

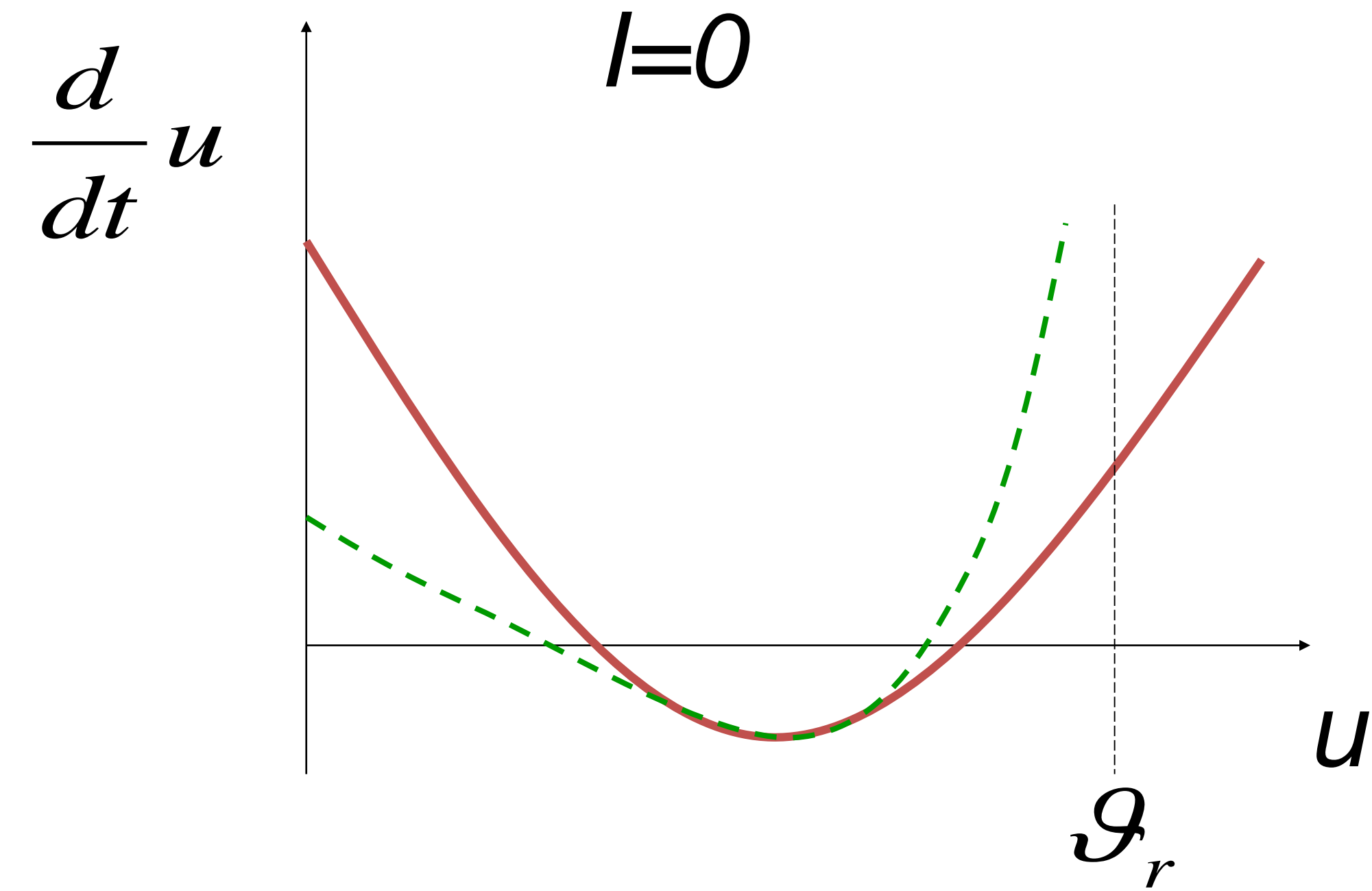
-Install PYTHON for Computer Exercises

-Exercise 3, on sheet

Videos (for today: 'week 1'):



Biological Modeling of Neural Networks – week1/Exercise 3



$$\tau \cdot \frac{d}{dt} u = F(u) + RI(t)$$

Homework!

Exercise 3: Integrate-and-fire models

The general form of an integrate-and-fire model is

$$\frac{du}{dt} = F(u) + \frac{RI(t)}{\tau} \quad (4)$$

where $F(u)$ is an appropriate function and $I(t)$ is the injected current. Three popular choices for the function F are the following (see Fig.1);

Leaky integrate-and-fire $F(u) = -\frac{u - u_{rest}}{\tau}$

Quadratic integrate-and-fire $F(u) = k \frac{(u - u_{rest})(u - u_{th})}{\tau}$

Exponential integrate-and-fire $F(u) = \frac{-(u - u_{rest}) + \Delta e^{\frac{u - u_{th}}{\Delta}}}{\tau}$.

3.1 Identify the resting potential u_{rest} and the spike threshold u_{th} in Fig. 1.

3.2 Consider three different values u_1, u_2 and u_3 for the voltage such that (i) u_1 is below u_{rest} (the resting potential), (ii) u_2 is between u_{rest} and u_{th} (the spike threshold), and (iii) u_3 is above u_{th} (see Fig. 1). For the three models described above, determine qualitatively the evolution of $u(t)$ when started at u_1, u_2 , and u_3 , assuming that the external input $I(t) \equiv 0$.

- For $u(t = 0) = u_1$, the voltage increases/decreases slowly/rapidly.
- For $u(t = 0) = u_2$,
- For $u(t = 0) = u_3$,

3.3 Why is u_{rest} called the resting potential? What is the role of u_{th} ?

3.4 Consider the two voltage traces shown in Fig. 2(b) (top) in response to a step current (bottom). Using the graphs in Fig. 2(a), determine which of the two models was used to generate each trace.

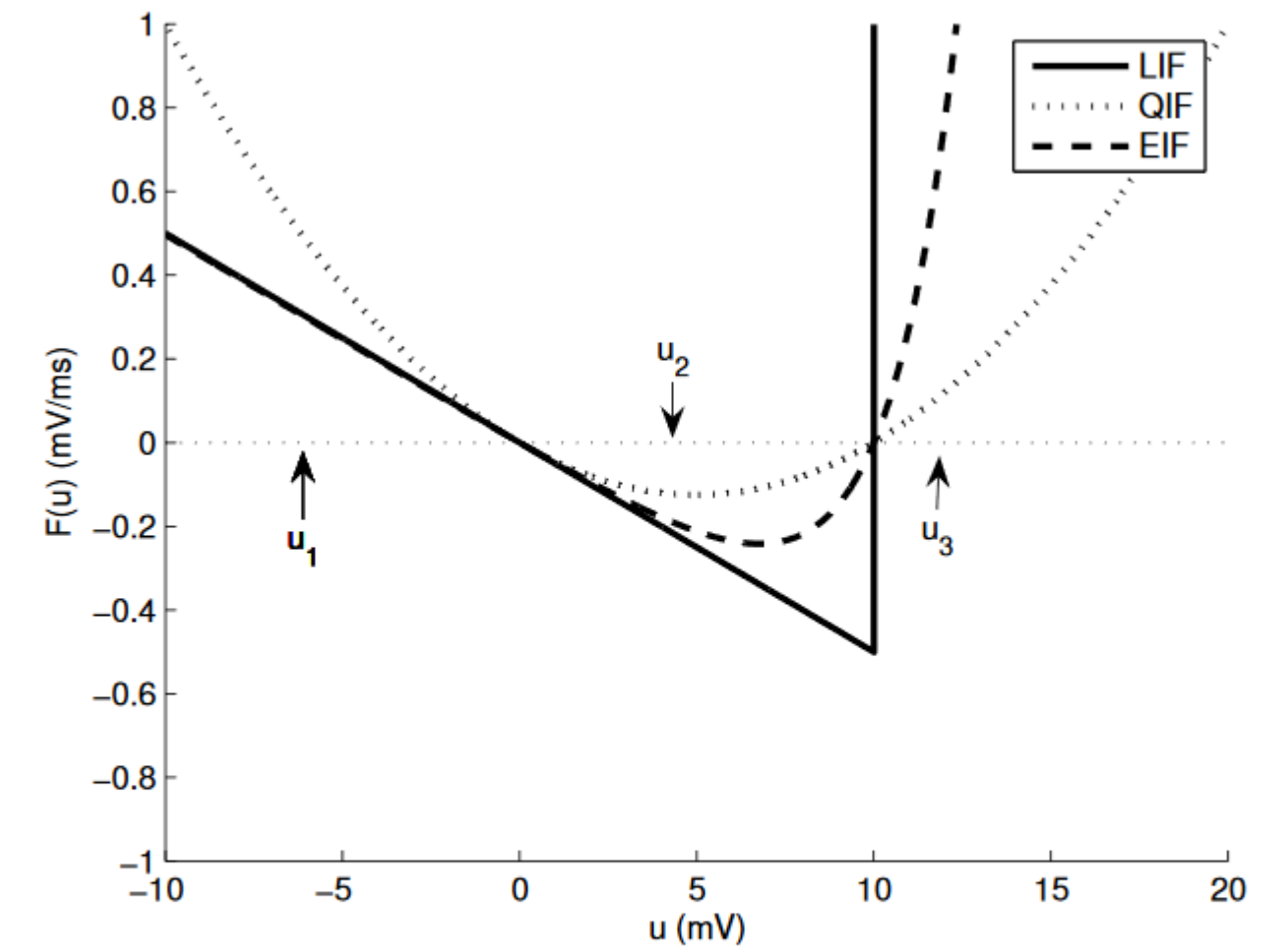


Figure 1: Sketch of the function $F(u)$ for three popular integrate-and-fire models.

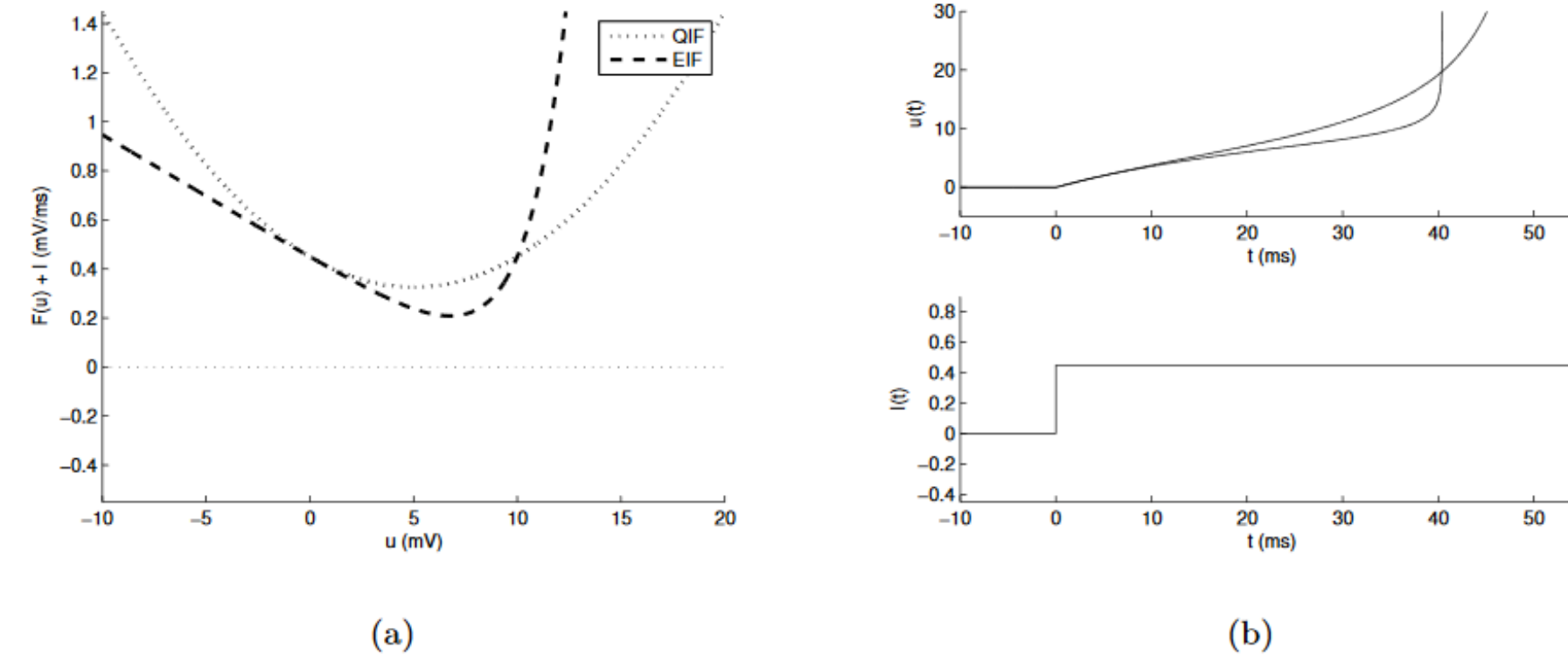


Figure 2: Left: Right-hand side of Eq. 4 for the quadratic and exponential integrate-and-fire models if a constant input current $I(t) > 0$ is applied. Lower right: Trace of the injected current. Upper right: Voltage trace of the two models (EIF and QIF).

Summary of Section 1.5.

The exponential integrate-and-fire (EIF) model is an excellent model of spike initiation in neurons. [- And better than the QIF which is 'too symmetric'].

The quality of a neuron model can be measured by predicting spikes for new input that was used to optimize the model parameters.

Further improvements of generalized integrate-and-fire models are possible and involve (i) noise in the spike generation process and (ii) adaptation.

First week – References and Suggested Reading

Reading: W. Gerstner, W.M. Kistler, R. Naud and L. Paninski, *Neuronal Dynamics: from single neurons to networks and models of cognition*. Chapter 1: Introduction. Cambridge Univ. Press, 2014

Selected references to linear and nonlinear integrate-and-fire models

- Lapicque, L. (1907). *Recherches quantitatives sur l'excitation électrique des nerfs traitée comme une polarisation*. J. Physiol. Pathol. Gen., 9:620-635.
- Stein, R. B. (1965). A theoretical analysis of neuronal variability. Biophys. J., 5:173-194.
- Ermentrout, G. B. (1996). *Type I membranes, phase resetting curves, and synchrony*. Neural Computation, 8(5):979-1001.
- Fourcaud-Trocme, N., Hansel, D., van Vreeswijk, C., and Brunel, N. (2003). *How spike generation mechanisms determine the neuronal response to fluctuating input*. J. Neuroscience, 23:11628-11640.
- Badel, L., Lefort, S., Berger, T., Petersen, C., Gerstner, W., and Richardson, M. (2008). Biological Cybernetics, 99(4-5):361-370.
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First week

THE END (of main lecture)

**MATH DETOUR SLIDES
(for online VIDEO)**

Week 1 – part 2: Detour/Linear differential equation



Neuronal Dynamics: Computational Neuroscience of Single Neurons

Week 1 – neurons and mathematics:
a first simple neuron model

Wulfram Gerstner

EPFL, Lausanne, Switzerland

√ 1.1 Neurons and Synapses:

Overview

√ 1.2 The Passive Membrane

- Linear circuit
- Dirac delta-function
- Detour: solution of 1-dim linear differential equation

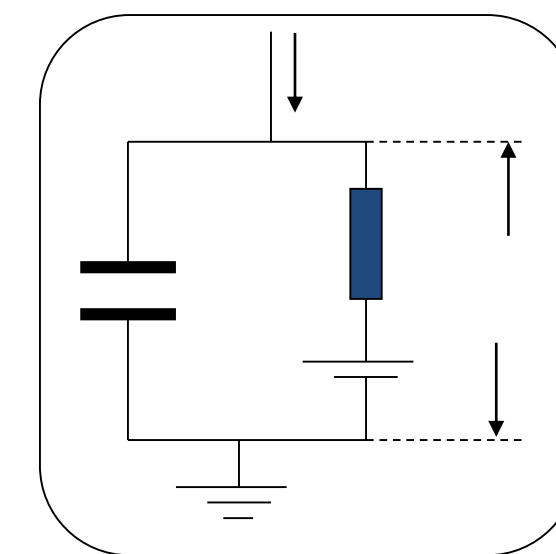
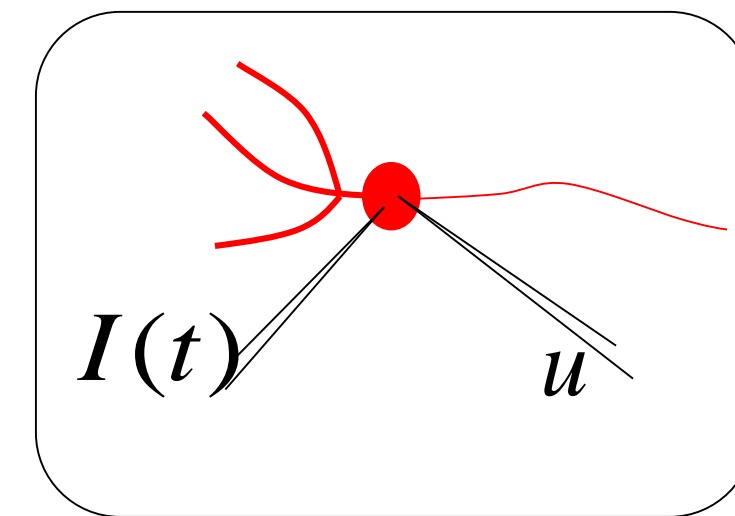
1.3 Leaky Integrate-and-Fire Model

1.4 Generalized Integrate-and-Fire Model

1.5. Quality of Integrate-and-Fire Models

Neuronal Dynamics – 1.2 Detour – Linear Differential Eq.

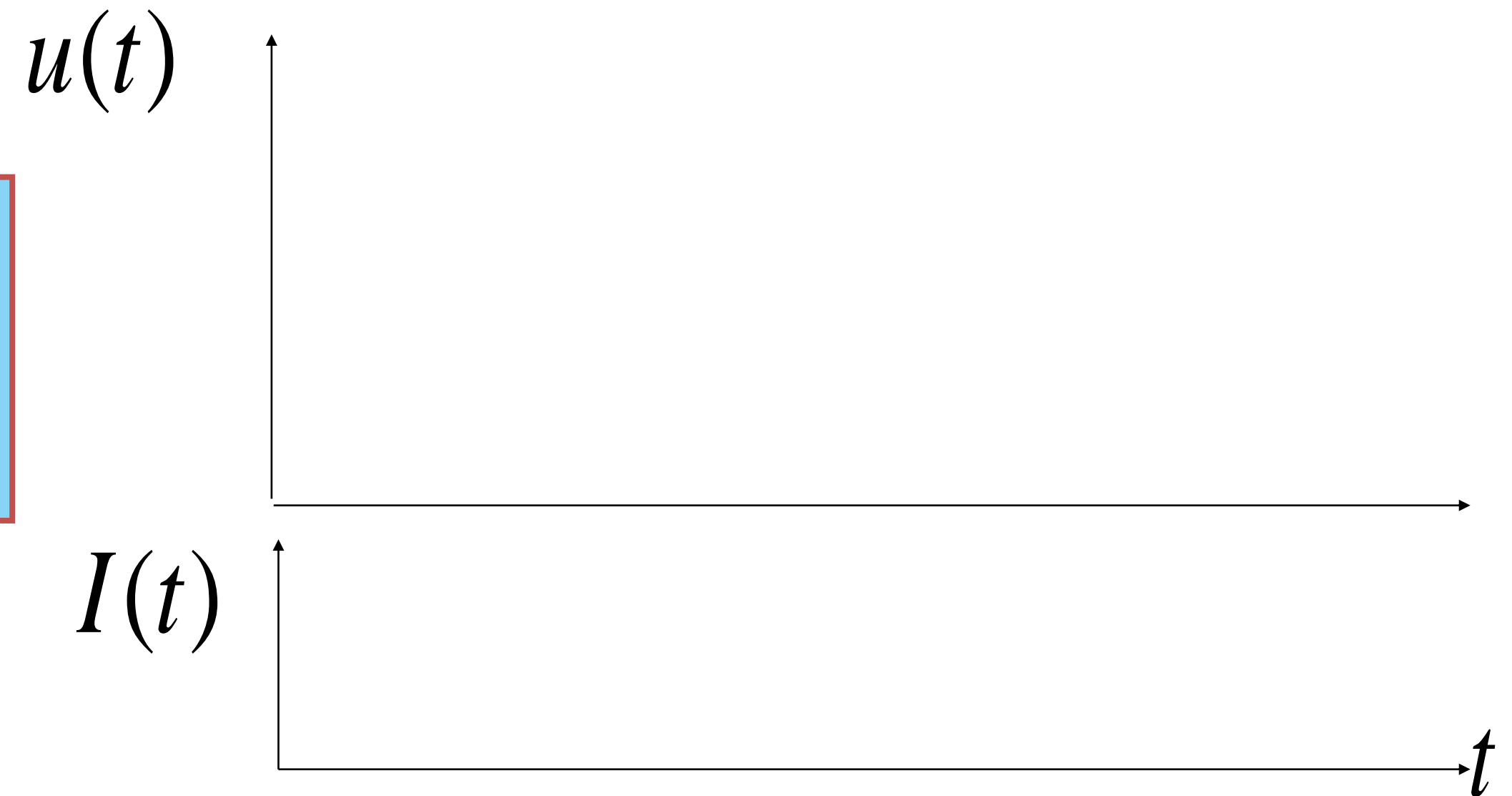
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$



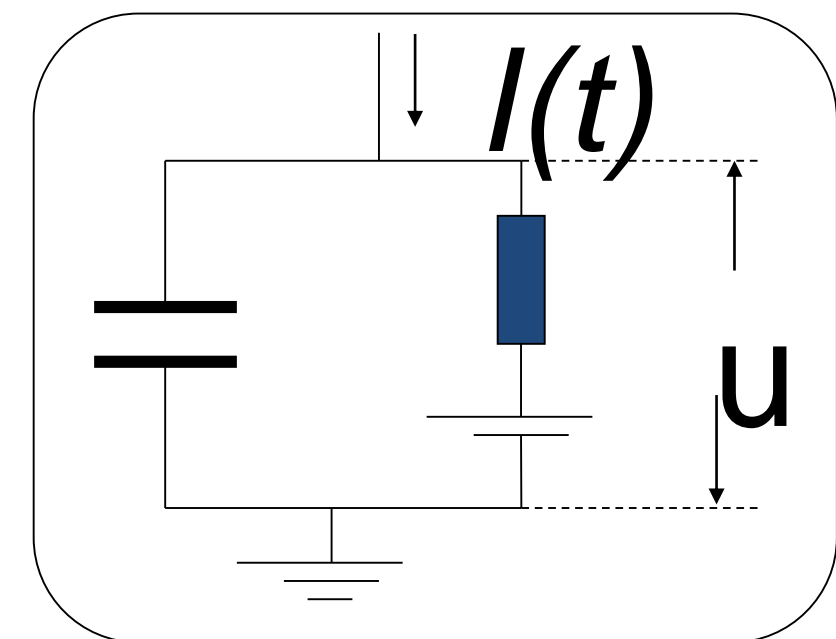
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Neuronal Dynamics – 1.2 Detour – Linear Differential Eq.

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

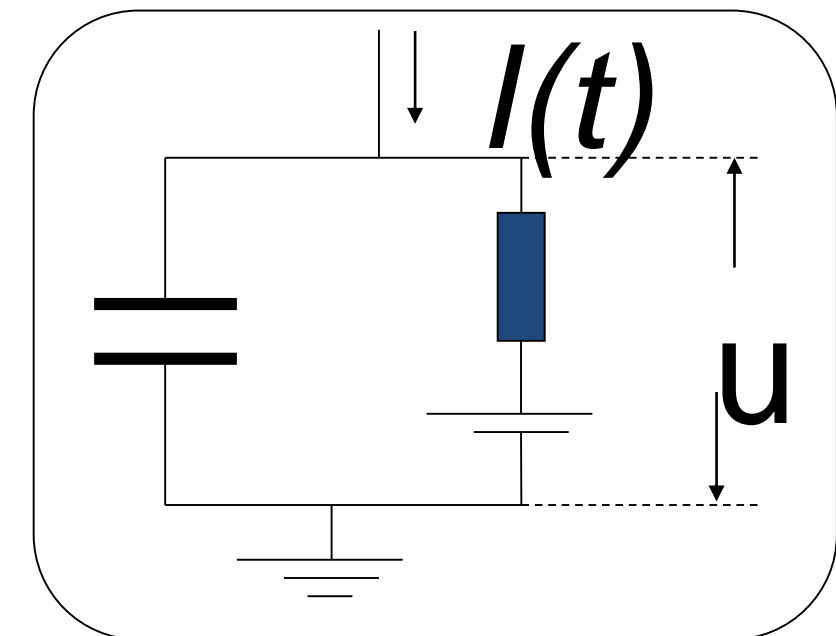
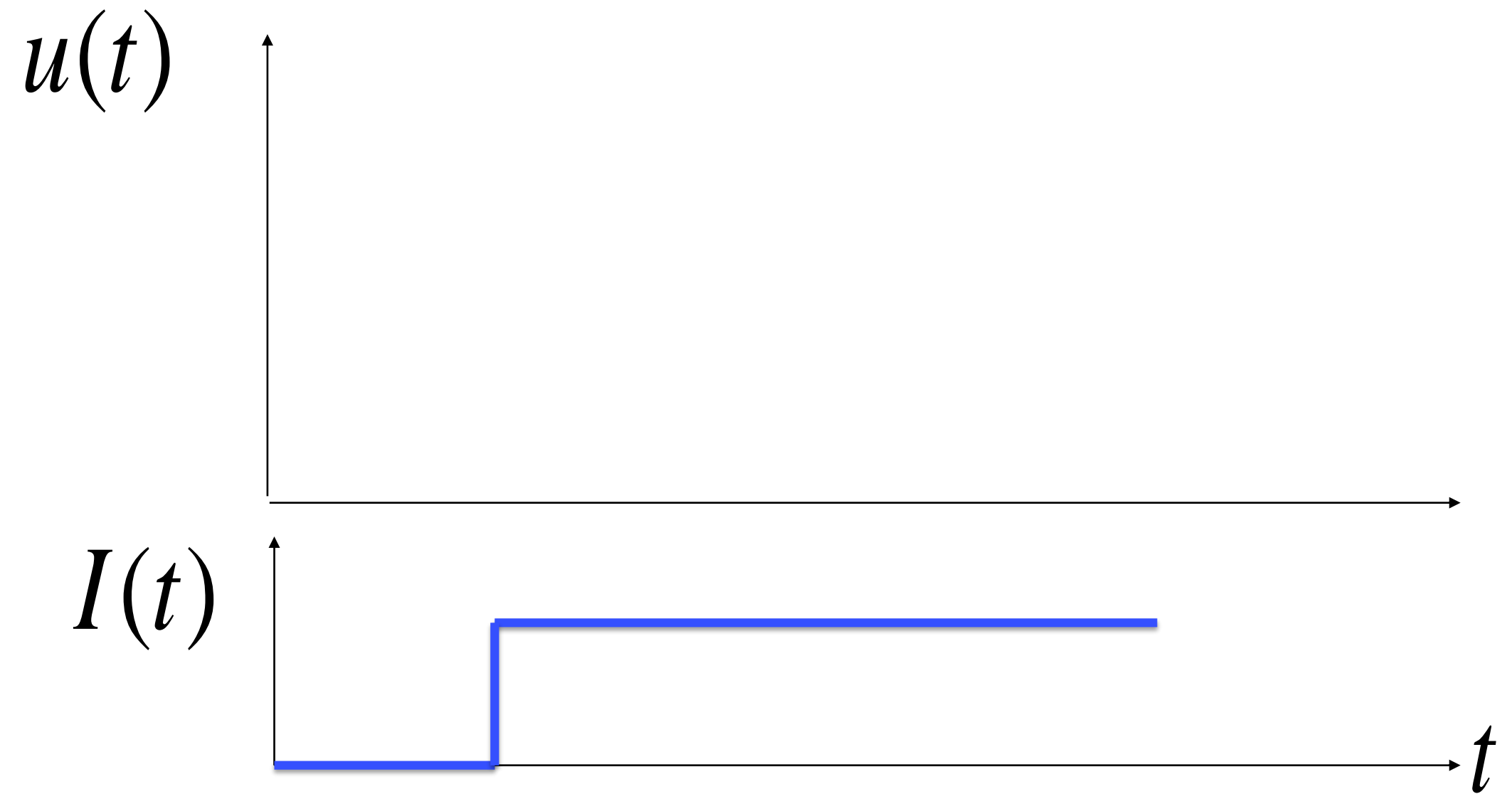


*Math development:
Response to step current*



Neuronal Dynamics – 1.2 Detour – Step current input

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$



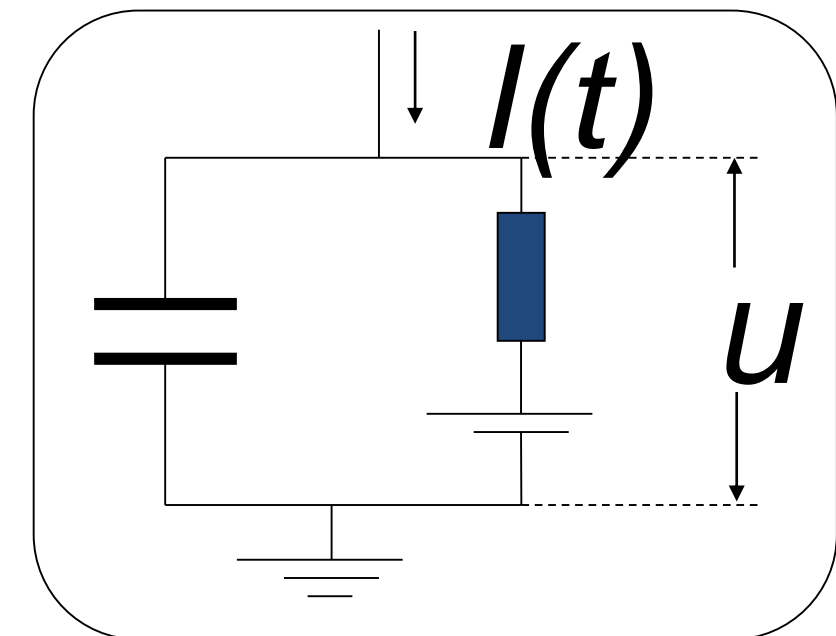
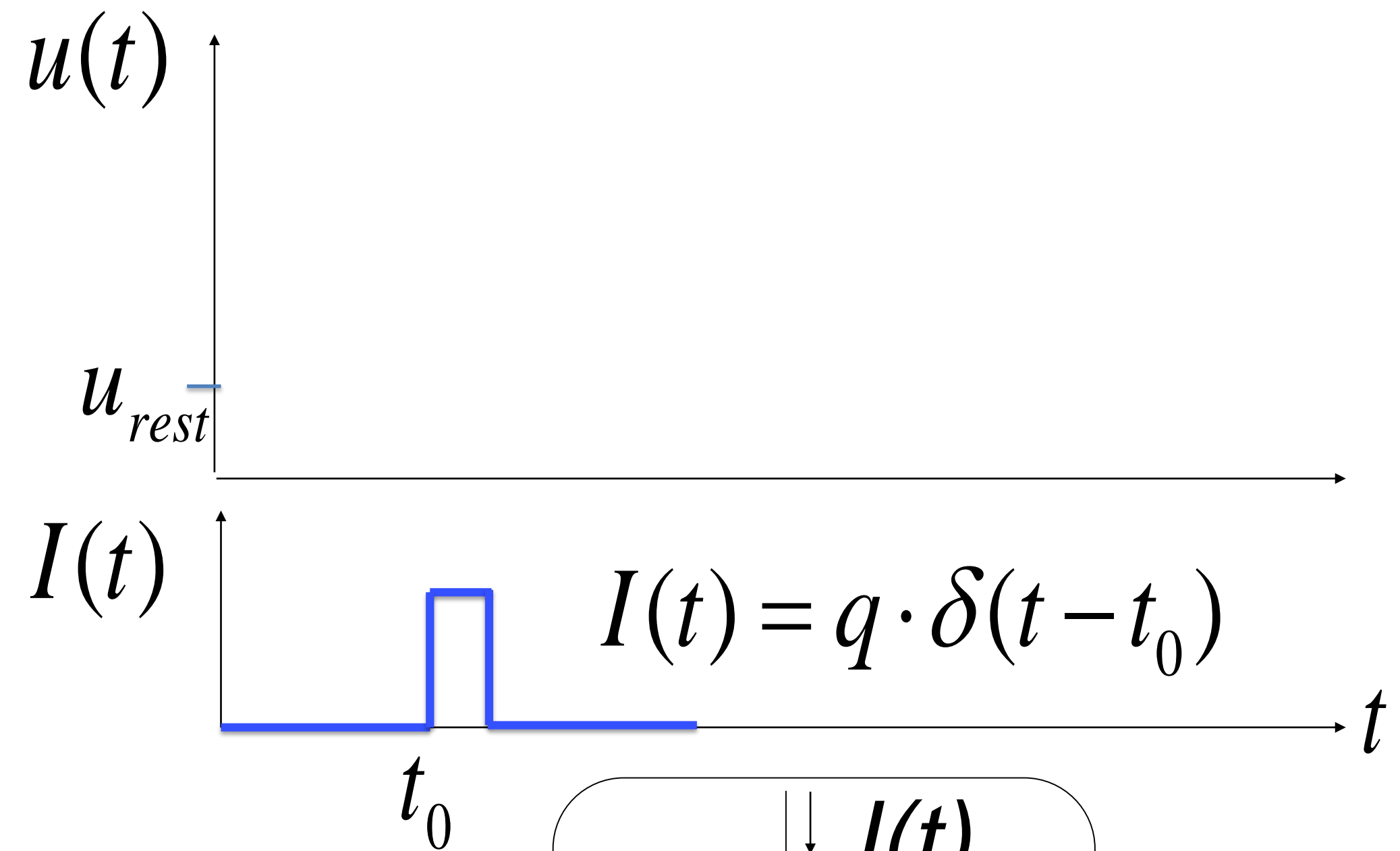
Neuronal Dynamics – 1.2 Detour – Short pulse input

$$u(t) = u_{rest} + RI_0 \left[1 - e^{-(t-t_0)/\tau} \right]$$

short pulse: $(t - t_0) \ll \tau$

*Math development:
Response to short
current pulse*

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

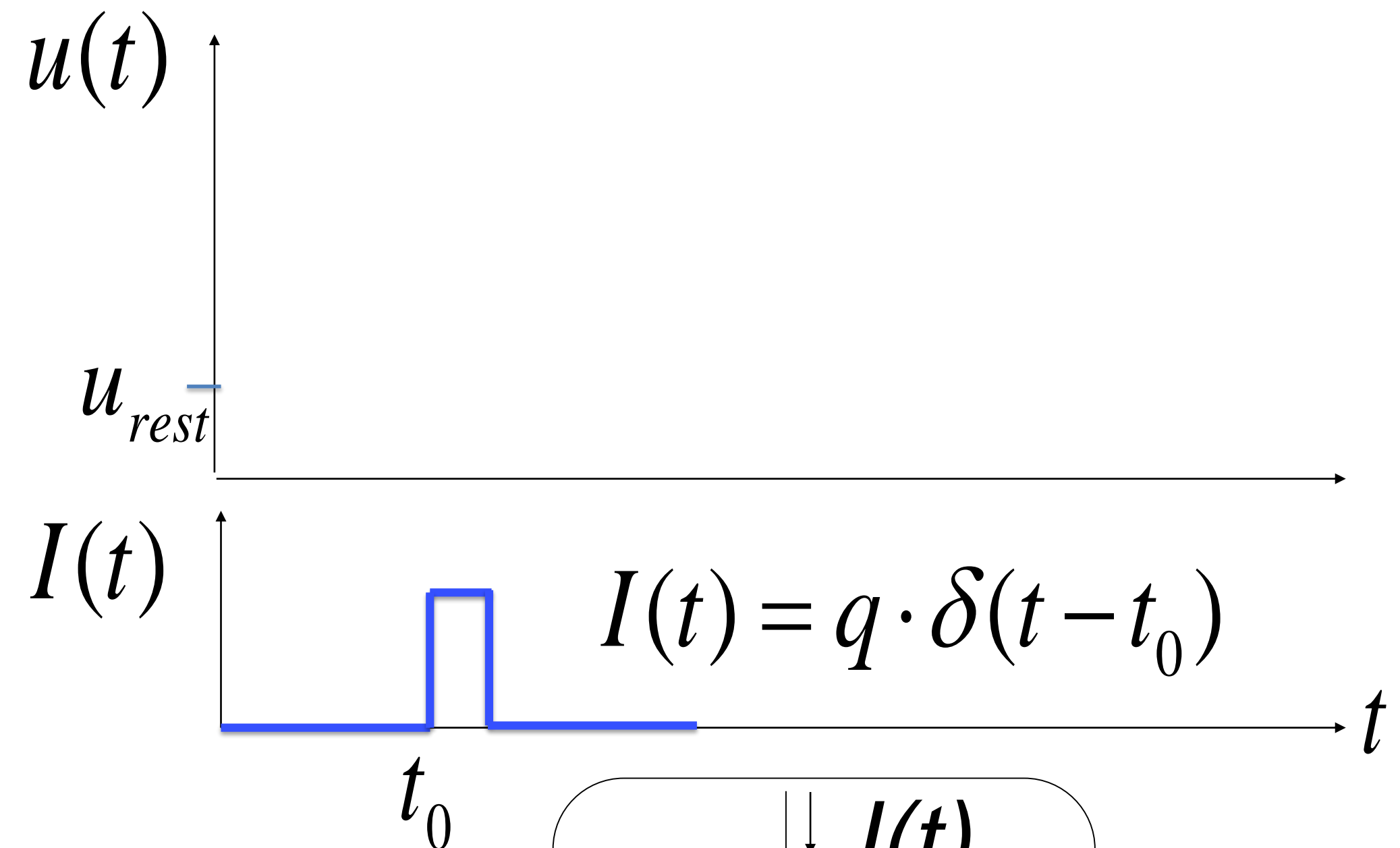


Neuronal Dynamics – 1.2 Detour – Short pulse input

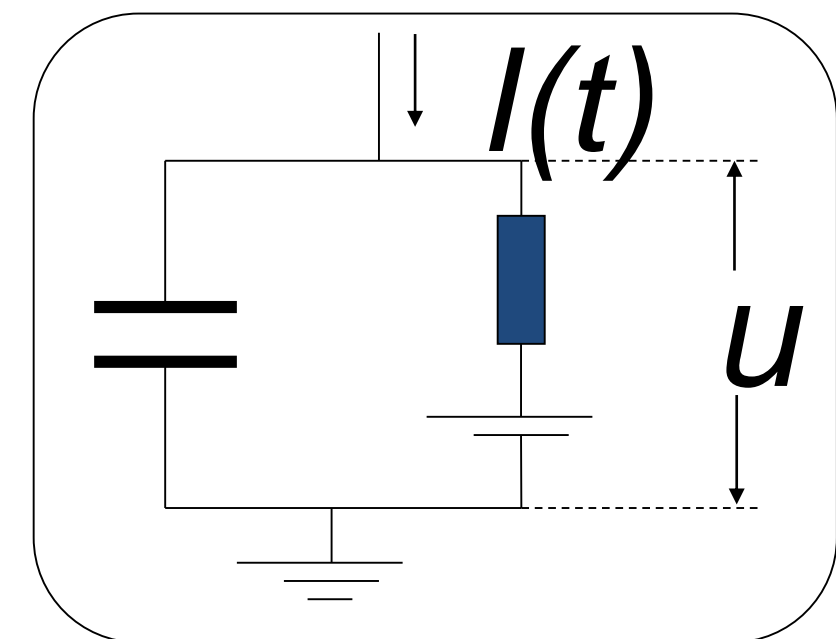
$$u(t) = u_{rest} + RI_0 \left[1 - e^{-(t-t_0)/\tau} \right]$$

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

short pulse: $(t - t_0) \ll \tau$



$$u(t) = u_{rest} + \frac{q}{C} e^{-(t-t_0)/\tau}$$



Neuronal Dynamics – 1.2 Detour – arbitrary input

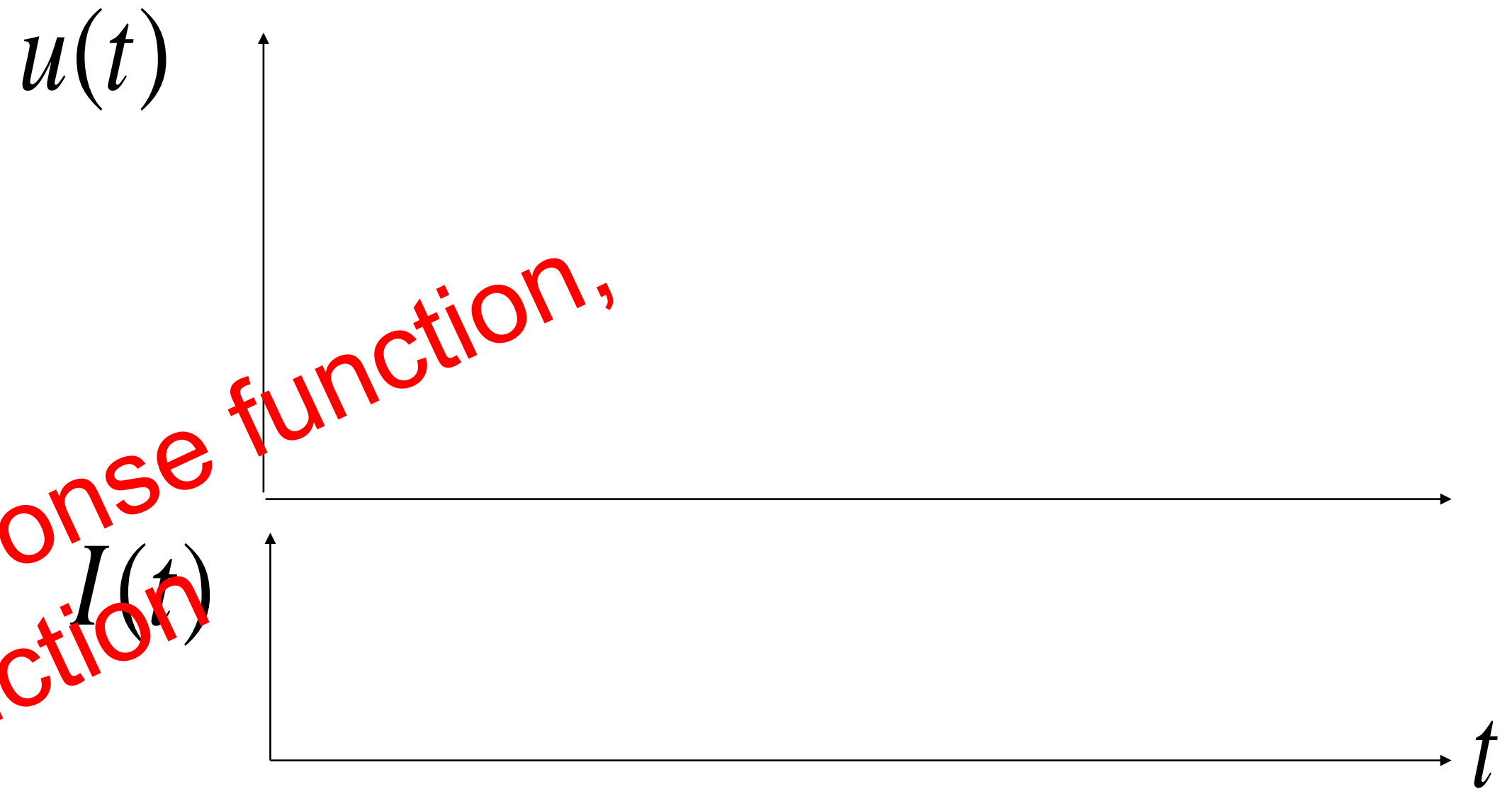
Single pulse

$$u(t) = u_{rest} + \frac{q}{C} e^{-(t-t_0)/\tau}$$

Multiple pulses:

$$u(t) = u_{rest} + \int_{-\infty}^t \frac{1}{C} e^{-(t-t')/\tau} I(t') dt'$$

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$



Neuronal Dynamics – 1.2 Detour – Greens function

Single pulse

$$\Delta u(t) = q \frac{1}{C} e^{-(t-t_0)/\tau}$$

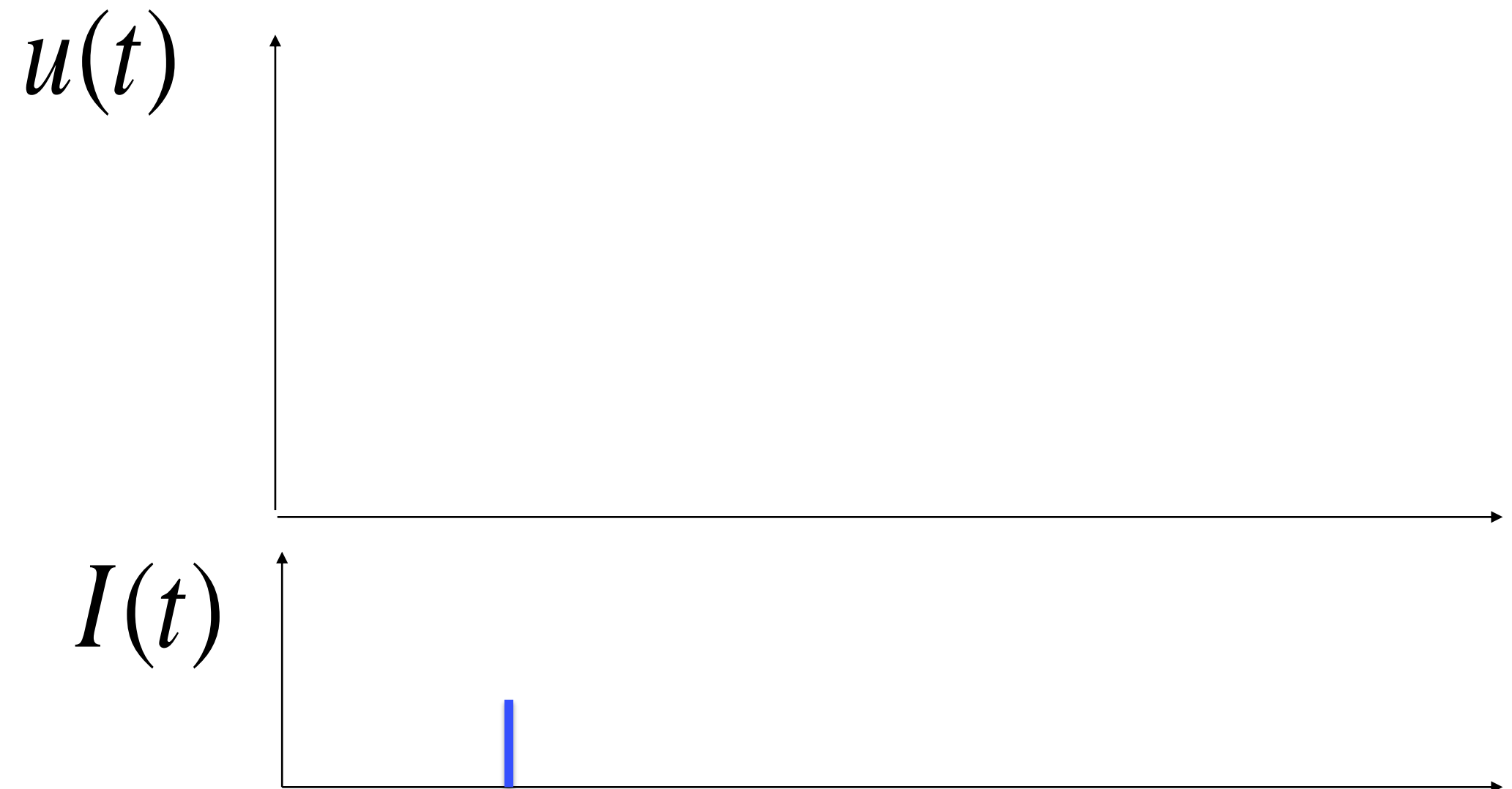
$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

Multiple pulses:

$$u(t) = u_{rest} + [u(t_0) - u_{rest}] + \int_{t_0}^t \frac{1}{C} e^{-(t-t')/\tau} I(t') dt'$$

Impulse response function,
Green's function

$$u(t) = u_{rest} + \int_{-\infty}^t \frac{1}{C} e^{-(t-t')/\tau} I(t') dt'$$



Neuronal Dynamics – 1.2 Detour – arbitrary input

$$\tau \cdot \frac{d}{dt} u = -(u - u_{rest}) + RI(t)$$

*If you don't feel at ease yet,
spend **10 minutes** on these
mathematical exercises
And quiz 2 in week 1.*

Arbitrary input

$$u(t) = u_{rest} + \int_{-\infty}^t \frac{1}{c} e^{-(t-t')/\tau} I(t') dt'$$

Single pulse

$$\Delta u(t) = \frac{q}{c} e^{-(t-t_0)/\tau}$$

*you need to know the solutions
of linear differential equations!*